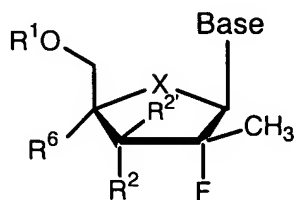


WE CLAIM:

1. A (2'R)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside ( $\beta$ -D or  $\beta$ -L) of the formula:



wherein

Base is a purine or pyrimidine base;

X is O, S, CH<sub>2</sub>, Se, NH, N-alkyl, CHW (R, S, or racemic), C(W)<sub>2</sub>, wherein W is F, Cl, Br, or I;

R<sup>1</sup> and R<sup>7</sup> are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid (or racemic mixture), a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein R<sup>1</sup> is H or phosphate; R<sup>2</sup> is H or phosphate; R<sup>1</sup> and R<sup>2</sup> or R<sup>7</sup> can also be linked with cyclic phosphate group;

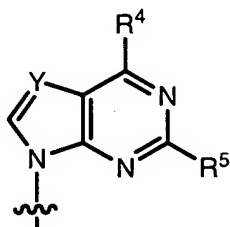
R<sup>2</sup> and R<sup>2'</sup> are independently H, C<sub>1-4</sub> alkyl, C<sub>1-4</sub> alkenyl, C<sub>1-4</sub> alkynyl, vinyl, N<sub>3</sub>, CN, Cl, Br, F, I, NO<sub>2</sub>, C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkenyl), C(O)O(C<sub>1-4</sub> alkynyl), C(O)O(C<sub>1-4</sub> alkenyl), O(C<sub>1-4</sub> acyl), O(C<sub>1-4</sub> alkyl), O(C<sub>1-4</sub> alkenyl), S(C<sub>1-4</sub> acyl), S(C<sub>1-4</sub> alkyl), S(C<sub>1-4</sub> alkynyl), S(C<sub>1-4</sub> alkenyl), SO(C<sub>1-4</sub> acyl), SO(C<sub>1-4</sub> alkyl), SO(C<sub>1-4</sub> alkynyl), SO(C<sub>1-4</sub> alkenyl),

SO<sub>2</sub>(C<sub>1-4</sub> acyl), SO<sub>2</sub>(C<sub>1-4</sub> alkyl), SO<sub>2</sub>(C<sub>1-4</sub> alkynyl), SO<sub>2</sub>(C<sub>1-4</sub> alkenyl),  
O<sub>3</sub>S(C<sub>1-4</sub> acyl), O<sub>3</sub>S(C<sub>1-4</sub> alkyl), O<sub>3</sub>S(C<sub>1-4</sub> alkenyl), NH<sub>2</sub>, NH(C<sub>1-4</sub>  
alkyl), NH(C<sub>1-4</sub> alkenyl), NH(C<sub>1-4</sub> alkynyl), NH(C<sub>1-4</sub> acyl), N(C<sub>1-4</sub>  
alkyl)<sub>2</sub>, N(C<sub>1-18</sub> acyl)<sub>2</sub>, wherein alkyl, alkynyl, alkenyl and vinyl are  
optionally substituted by N<sub>3</sub>, CN, one to three halogen (Cl, Br, F, I),  
NO<sub>2</sub>, C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkynyl),  
C(O)O(C<sub>1-4</sub> alkenyl), O(C<sub>1-4</sub> acyl), O(C<sub>1-4</sub> alkyl), O(C<sub>1-4</sub> alkenyl), S(C<sub>1-4</sub>  
acyl), S(C<sub>1-4</sub> alkyl), S(C<sub>1-4</sub> alkynyl), S(C<sub>1-4</sub> alkenyl), SO(C<sub>1-4</sub> acyl),  
SO(C<sub>1-4</sub> alkyl), SO(C<sub>1-4</sub> alkynyl), SO(C<sub>1-4</sub> alkenyl), SO<sub>2</sub>(C<sub>1-4</sub> acyl),  
SO<sub>2</sub>(C<sub>1-4</sub> alkyl), SO<sub>2</sub>(C<sub>1-4</sub> alkynyl), SO<sub>2</sub>(C<sub>1-4</sub> alkenyl), O<sub>3</sub>S(C<sub>1-4</sub> acyl),  
O<sub>3</sub>S(C<sub>1-4</sub> alkyl), O<sub>3</sub>S(C<sub>1-4</sub> alkenyl), NH<sub>2</sub>, NH(C<sub>1-4</sub> alkyl), NH(C<sub>1-4</sub>  
alkenyl), NH(C<sub>1-4</sub> alkynyl), NH(C<sub>1-4</sub> acyl), N(C<sub>1-4</sub> alkyl)<sub>2</sub>, N(C<sub>1-4</sub>  
acyl)<sub>2</sub>, OR<sup>7</sup>; R<sup>2</sup> and R<sup>2'</sup> can be linked together to form a vinyl optionally  
substituted by one or two of N<sub>3</sub>, CN, Cl, Br, F, I, NO<sub>2</sub>; and

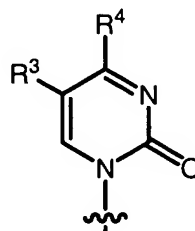
R<sup>6</sup> is an optionally substituted alkyl (including lower alkyl), cyano (CN), CH<sub>3</sub>,  
OCH<sub>3</sub>, OCH<sub>2</sub>CH<sub>3</sub>, hydroxy methyl (CH<sub>2</sub>OH), fluoromethyl (CH<sub>2</sub>F),  
azido (N<sub>3</sub>), CHCN, CH<sub>2</sub>N<sub>3</sub>, CH<sub>2</sub>NH<sub>2</sub>, CH<sub>2</sub>NHCH<sub>3</sub>, CH<sub>2</sub>N(CH<sub>3</sub>)<sub>2</sub>,  
alkyne (optionally substituted), or fluoro;

or its pharmaceutically acceptable salt or prodrug thereof.

2. The (2'*R*)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside (β-D or β-L) of claim 1 or  
its pharmaceutically acceptable salt or prodrug thereof, wherein Base is selected from the  
group consisting of:



(a)



(b)

wherein

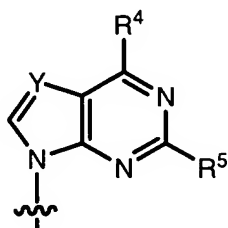
Y is N or CH.

$R^3$ ,  $R^4$  and  $R^5$  are independently H, halogen including F, Cl, Br, I, OH, OR', SH, SR', NH<sub>2</sub>, NHR', NR'<sub>2</sub>, lower alkyl of C<sub>1</sub>-C<sub>6</sub>, halogenated (F, Cl, Br, I) lower alkyl of C<sub>1</sub>-C<sub>6</sub> such as CF<sub>3</sub> and CH<sub>2</sub>CH<sub>2</sub>F, lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CH<sub>2</sub>, halogenated (F, Cl, Br, I) lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CHCl, CH=CHBr and CH=CHI, lower alkynyl of C<sub>2</sub>-C<sub>6</sub> such as C≡CH, halogenated (F, Cl, Br, I) lower alkynyl of C<sub>2</sub>-C<sub>6</sub>, lower alkoxy of C<sub>1</sub>-C<sub>6</sub> such as CH<sub>2</sub>OH and CH<sub>2</sub>CH<sub>2</sub>OH, halogenated (F, Cl, Br, I) lower alkoxy of C<sub>1</sub>-C<sub>6</sub>, CO<sub>2</sub>H, CO<sub>2</sub>R', CONH<sub>2</sub>, CONHR', CONR'<sub>2</sub>, CH=CHCO<sub>2</sub>H, CH=CHCO<sub>2</sub>R'; and,

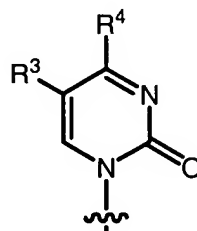
R' is an optionally substituted alkyl of C<sub>1</sub>-C<sub>12</sub> (particularly when the alkyl is an amino acid residue), cycloalkyl, optionally substituted alkynyl of C<sub>2</sub>-C<sub>6</sub>, optionally substituted lower alkenyl of C<sub>2</sub>-C<sub>6</sub>, or optionally substituted acyl.

3. The (2'R)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside (β-D) of claim 1 or its pharmaceutically acceptable salt or prodrug thereof,

wherein Base is selected from the group consisting of (a) or (b):



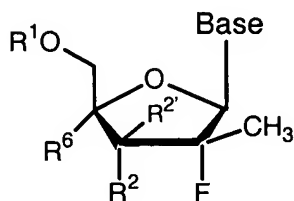
(a)



(b)

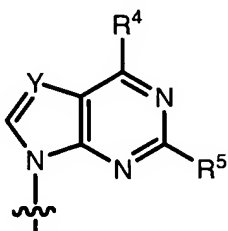
and wherein R<sup>1</sup> is H, R<sup>2</sup> is OH, R<sup>2'</sup> is H, R<sup>3</sup> is H, and R<sup>4</sup> is NH<sub>2</sub> or OH, and R<sup>5</sup> is NH<sub>2</sub>.

4. A (2'R)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside ( $\beta$ -D or  $\beta$ -L) of the formula:

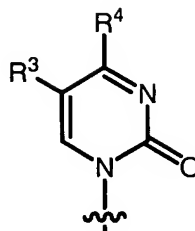


wherein

- 5 Base is selected from the group consisting of



(a)



(b)

Y is N or CH;

- 10  $R^1$  and  $R^7$  are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including
- 15 methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid (or racemic mixture), a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein  $R^1$  is H or phosphate;  $R^2$  is H or phosphate;  $R^1$  and  $R^2$  or  $R^7$  can also be linked with cyclic phosphate group;
- 20



$R^2$  and  $R^{2'}$  are independently H,  $C_{1-4}$  alkyl,  $C_{1-4}$  alkenyl,  $C_{1-4}$  alkynyl, vinyl,  $N_3$ ,  
 CN, Cl, Br, F, I,  $NO_2$ ,  $C(O)O(C_{1-4} \text{ alkyl})$ ,  $C(O)O(C_{1-4} \text{ alkyl})$ ,  $C(O)O(C_{1-4}$   
 $C_{1-4} \text{ alkynyl})$ ,  $C(O)O(C_{1-4} \text{ alkenyl})$ ,  $O(C_{1-4} \text{ acyl})$ ,  $O(C_{1-4} \text{ alkyl})$ ,  $O(C_{1-4}$   
 $\text{alkenyl})$ ,  $S(C_{1-4} \text{ acyl})$ ,  $S(C_{1-4} \text{ alkyl})$ ,  $S(C_{1-4} \text{ alkynyl})$ ,  $S(C_{1-4} \text{ alkenyl})$ ,  
 5  $SO(C_{1-4} \text{ acyl})$ ,  $SO(C_{1-4} \text{ alkyl})$ ,  $SO(C_{1-4} \text{ alkynyl})$ ,  $SO(C_{1-4} \text{ alkenyl})$ ,  
 $SO_2(C_{1-4} \text{ acyl})$ ,  $SO_2(C_{1-4} \text{ alkyl})$ ,  $SO_2(C_{1-4} \text{ alkynyl})$ ,  $SO_2(C_{1-4} \text{ alkenyl})$ ,  
 $O_3S(C_{1-4} \text{ acyl})$ ,  $O_3S(C_{1-4} \text{ alkyl})$ ,  $O_3S(C_{1-4} \text{ alkenyl})$ ,  $NH_2$ ,  $NH(C_{1-4}$   
 $\text{alkyl})$ ,  $NH(C_{1-4} \text{ alkenyl})$ ,  $NH(C_{1-4} \text{ alkynyl})$ ,  $NH(C_{1-4} \text{ acyl})$ ,  $N(C_{1-4}$   
 $\text{alkyl})_2$ ,  $N(C_{1-18} \text{ acyl})_2$ , wherein alkyl, alkynyl, alkenyl and vinyl are  
 10 optionally substituted by  $N_3$ , CN, one to three halogen (Cl, Br, F, I),  
 $NO_2$ ,  $C(O)O(C_{1-4} \text{ alkyl})$ ,  $C(O)O(C_{1-4} \text{ alkyl})$ ,  $C(O)O(C_{1-4} \text{ alkynyl})$ ,  
 $C(O)O(C_{1-4} \text{ alkenyl})$ ,  $O(C_{1-4} \text{ acyl})$ ,  $O(C_{1-4} \text{ alkyl})$ ,  $O(C_{1-4} \text{ alkenyl})$ ,  $S(C_{1-4}$   
 $\text{acyl})$ ,  $S(C_{1-4} \text{ alkyl})$ ,  $S(C_{1-4} \text{ alkynyl})$ ,  $S(C_{1-4} \text{ alkenyl})$ ,  $SO(C_{1-4} \text{ acyl})$ ,  
 $SO(C_{1-4} \text{ alkyl})$ ,  $SO(C_{1-4} \text{ alkynyl})$ ,  $SO(C_{1-4} \text{ alkenyl})$ ,  $SO_2(C_{1-4} \text{ acyl})$ ,  
 15  $SO_2(C_{1-4} \text{ alkyl})$ ,  $SO_2(C_{1-4} \text{ alkynyl})$ ,  $SO_2(C_{1-4} \text{ alkenyl})$ ,  $O_3S(C_{1-4} \text{ acyl})$ ,  
 $O_3S(C_{1-4} \text{ alkyl})$ ,  $O_3S(C_{1-4} \text{ alkenyl})$ ,  $NH_2$ ,  $NH(C_{1-4} \text{ alkyl})$ ,  $NH(C_{1-4}$   
 $\text{alkenyl})$ ,  $NH(C_{1-4} \text{ alkynyl})$ ,  $NH(C_{1-4} \text{ acyl})$ ,  $N(C_{1-4} \text{ alkyl})_2$ ,  $N(C_{1-4}$   
 $\text{acyl})_2$ ,  $OR^7$ ;  $R^2$  and  $R^{2'}$  can be linked together to form a vinyl optionally  
 substituted by one or two of  $N_3$ , CN, Cl, Br, F, I,  $NO_2$ ;

$R^3$ ,  $R^4$  and  $R^5$  are independently H, halogen including F, Cl, Br, I, OH,  $OR'$ ,  
 SH,  $SR'$ ,  $NH_2$ ,  $NHR'$ ,  $NR'_2$ , lower alkyl of  $C_1-C_6$ , halogenated (F, Cl,  
 Br, I) lower alkyl of  $C_1-C_6$  such as  $CF_3$  and  $CH_2CH_2F$ , lower alkenyl of  
 $C_2-C_6$  such as  $CH=CH_2$ , halogenated (F, Cl, Br, I) lower alkenyl of  $C_2-$   
 $C_6$  such as  $CH=CHCl$ ,  $CH=CHBr$  and  $CH=CHI$ , lower alkynyl of  $C_2-$   
 25  $C_6$  such as  $C\equiv CH$ , halogenated (F, Cl, Br, I) lower alkynyl of  $C_2-C_6$ ,  
 lower alkoxy of  $C_1-C_6$  such as  $CH_2OH$  and  $CH_2CH_2OH$ , halogenated  
 (F, Cl, Br, I) lower alkoxy of  $C_1-C_6$ ,  $CO_2H$ ,  $CO_2R'$ ,  $CONH_2$ ,  $CONHR'$ ,  
 $CONR'_2$ ,  $CH=CHCO_2H$ ,  $CH=CHCO_2R'$ ;

$R'$  is an optionally substituted alkyl of  $C_1-C_{12}$  (particularly when the alkyl is an  
 30 amino acid residue), cycloalkyl, optionally substituted alkynyl of  $C_2-$

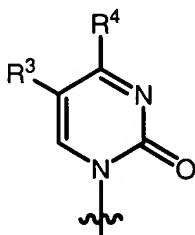
C<sub>6</sub>, optionally substituted lower alkenyl of C<sub>2</sub>-C<sub>6</sub>, or optionally substituted acyl;

R<sup>6</sup> is an optionally substituted alkyl (including lower alkyl), cyano (CN), CH<sub>3</sub>, OCH<sub>3</sub>, OCH<sub>2</sub>CH<sub>3</sub>, hydroxy methyl (CH<sub>2</sub>OH), fluoromethyl (CH<sub>2</sub>F), azido (N<sub>3</sub>), CHCN, CH<sub>2</sub>N<sub>3</sub>, CH<sub>2</sub>NH<sub>2</sub>, CH<sub>2</sub>NHCH<sub>3</sub>, CH<sub>2</sub>N(CH<sub>3</sub>)<sub>2</sub>, alkyne (optionally substituted), or fluoro;

or its pharmaceutically acceptable salt or prodrug thereof.

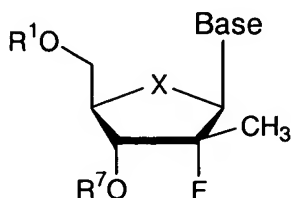
5. The (2'*R*)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside (β-D) of claim 4 or its pharmaceutically acceptable salt or prodrug thereof, wherein

Base is



and R<sup>1</sup> is H, R<sup>2</sup> is OH, R<sup>2'</sup> is H, R<sup>3</sup> is H, R<sup>4</sup> is NH<sub>2</sub> or OH, and R<sup>6</sup> is H.

6. A (2'*R*)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside (β-D or β-L) or its pharmaceutically acceptable salt or prodrug thereof of the structure:



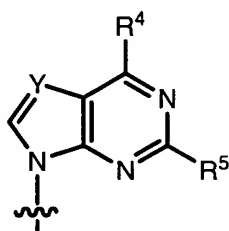
wherein Base is a purine or pyrimidine base;

X is O, S, CH<sub>2</sub>, Se, NH, N-alkyl, CHW (R, S, or racemic), C(W)<sub>2</sub>, wherein W is F, Cl, Br, or I; and,

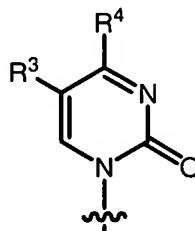
R<sup>1</sup> and R<sup>7</sup> are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid, a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein R<sup>1</sup> or R<sup>7</sup> is independently H or phosphate; R<sup>1</sup> and R<sup>7</sup> can also be linked with cyclic phosphate group.

7. The (2'*R*)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside (β-D or β-L) of claim 6 or its pharmaceutically acceptable salt or prodrug thereof,

wherein Base is selected from the group consisting of:



(a)



(b)

Y is N or CH;

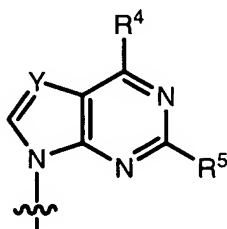
R<sup>3</sup>, R<sup>4</sup> and R<sup>5</sup> are independently H, halogen including F, Cl, Br, I, OH, OR', SH, SR', NH<sub>2</sub>, NHR', NR'<sub>2</sub>, lower alkyl of C<sub>1</sub>-C<sub>6</sub>, halogenated (F, Cl,

Br, I) lower alkyl of C<sub>1</sub>-C<sub>6</sub> such as CF<sub>3</sub> and CH<sub>2</sub>CH<sub>2</sub>F, lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CH<sub>2</sub>, halogenated (F, Cl, Br, I) lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CHCl, CH=CHBr and CH=CHI, lower alkynyl of C<sub>2</sub>-C<sub>6</sub> such as C≡CH, halogenated (F, Cl, Br, I) lower alkynyl of C<sub>2</sub>-C<sub>6</sub>, lower alkoxy of C<sub>1</sub>-C<sub>6</sub> such as CH<sub>2</sub>OH and CH<sub>2</sub>CH<sub>2</sub>OH, halogenated (F, Cl, Br, I) lower alkoxy of C<sub>1</sub>-C<sub>6</sub>, CO<sub>2</sub>H, CO<sub>2</sub>R', CONH<sub>2</sub>, CONHR', CONR'<sub>2</sub>, CH=CHCO<sub>2</sub>H, CH=CHCO<sub>2</sub>R'; and,

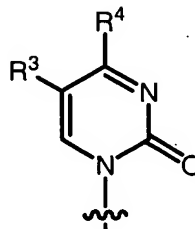
R' is an optionally substituted alkyl of C<sub>1</sub>-C<sub>12</sub> (particularly when the alkyl is an amino acid residue), cycloalkyl, optionally substituted alkynyl of C<sub>2</sub>-C<sub>6</sub>, optionally substituted lower alkenyl of C<sub>2</sub>-C<sub>6</sub>, or optionally substituted acyl.

8. The (2'*R*)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside (β-D) of claim 6 or its pharmaceutically acceptable salt or prodrug thereof,

wherein Base is selected from the group consisting of (a) or (b):



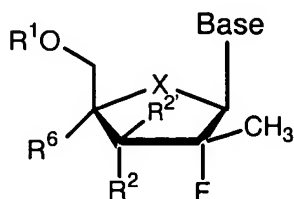
(a)



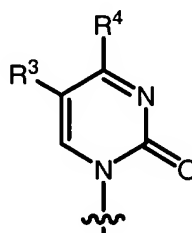
(b)

and wherein R<sup>1</sup> and R<sup>7</sup> are H, R<sup>3</sup> is H, and R<sup>4</sup> is NH<sub>2</sub> or OH, and R<sup>5</sup> is NH<sub>2</sub>.

9. A (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside ( $\beta$ -D or  $\beta$ -L) of the formula:



wherein Base is



5 X is O, S, CH<sub>2</sub>, Se, NH, N-alkyl, CHW (R, S, or racemic), C(W)<sub>2</sub>, wherein W is F, Cl, Br, or I;

R<sup>1</sup> and R<sup>7</sup> are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including  
 10 optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid (or  
 15 racemic mixture), a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein R<sup>1</sup> is H or phosphate; R<sup>2</sup> is H or phosphate; R<sup>1</sup> and R<sup>2</sup> or R<sup>7</sup> can also be linked with cyclic phosphate group;

20 R<sup>2</sup> and R<sup>2'</sup> are independently H, C<sub>1-4</sub> alkyl, C<sub>1-4</sub> alkenyl, C<sub>1-4</sub> alkynyl, vinyl, N<sub>3</sub>, CN, Cl, Br, F, I, NO<sub>2</sub>, C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkynyl), C(O)O(C<sub>1-4</sub> alkenyl), O(C<sub>1-4</sub> acyl), O(C<sub>1-4</sub> alkyl), O(C<sub>1-4</sub>

5 alkenyl), S(C<sub>1-4</sub> acyl), S(C<sub>1-4</sub> alkyl), S(C<sub>1-4</sub> alkynyl), S(C<sub>1-4</sub> alkenyl),  
 SO(C<sub>1-4</sub> acyl), SO(C<sub>1-4</sub> alkyl), SO(C<sub>1-4</sub> alkynyl), SO(C<sub>1-4</sub> alkenyl),  
 SO<sub>2</sub>(C<sub>1-4</sub> acyl), SO<sub>2</sub>(C<sub>1-4</sub> alkyl), SO<sub>2</sub>(C<sub>1-4</sub> alkynyl), SO<sub>2</sub>(C<sub>1-4</sub> alkenyl),  
 O<sub>3</sub>S(C<sub>1-4</sub> acyl), O<sub>3</sub>S(C<sub>1-4</sub> alkyl), O<sub>3</sub>S(C<sub>1-4</sub> alkenyl), NH<sub>2</sub>, NH(C<sub>1-4</sub>  
 10 alkyl), NH(C<sub>1-4</sub> alkenyl), NH(C<sub>1-4</sub> alkynyl), NH(C<sub>1-4</sub> acyl), N(C<sub>1-4</sub>  
 alkyl)<sub>2</sub>, N(C<sub>1-18</sub> acyl)<sub>2</sub>, wherein alkyl, alkynyl, alkenyl and vinyl are  
 optionally substituted by N<sub>3</sub>, CN, one to three halogen (Cl, Br, F, I),  
 NO<sub>2</sub>, C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkynyl),  
 C(O)O(C<sub>1-4</sub> alkenyl), O(C<sub>1-4</sub> acyl), O(C<sub>1-4</sub> alkyl), O(C<sub>1-4</sub> alkenyl), S(C<sub>1-4</sub>  
 15 4 acyl), S(C<sub>1-4</sub> alkyl), S(C<sub>1-4</sub> alkynyl), S(C<sub>1-4</sub> alkenyl), SO(C<sub>1-4</sub> acyl),  
 SO(C<sub>1-4</sub> alkyl), SO(C<sub>1-4</sub> alkynyl), SO(C<sub>1-4</sub> alkenyl), SO<sub>2</sub>(C<sub>1-4</sub> acyl),  
 SO<sub>2</sub>(C<sub>1-4</sub> alkyl), SO<sub>2</sub>(C<sub>1-4</sub> alkynyl), SO<sub>2</sub>(C<sub>1-4</sub> alkenyl), O<sub>3</sub>S(C<sub>1-4</sub> acyl),  
 O<sub>3</sub>S(C<sub>1-4</sub> alkyl), O<sub>3</sub>S(C<sub>1-4</sub> alkenyl), NH<sub>2</sub>, NH(C<sub>1-4</sub> alkyl), NH(C<sub>1-4</sub>  
 alkenyl), NH(C<sub>1-4</sub> alkynyl), NH(C<sub>1-4</sub> acyl), N(C<sub>1-4</sub> alkyl)<sub>2</sub>, N(C<sub>1-4</sub>  
 20 acyl)<sub>2</sub>, OR<sup>7</sup>; R<sup>2</sup> and R<sup>2'</sup> can be linked together to form a vinyl optionally  
 substituted by one or two of N<sub>3</sub>, CN, Cl, Br, F, I, NO<sub>2</sub>;

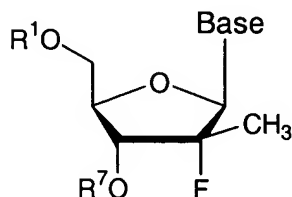
R<sup>3</sup> and R<sup>4</sup> are independently H, halogen including F, Cl, Br, I, OH, OR', SH,  
 SR', NH<sub>2</sub>, NHR', NR'<sub>2</sub>, lower alkyl of C<sub>1</sub>-C<sub>6</sub>, halogenated (F, Cl, Br, I)  
 lower alkyl of C<sub>1</sub>-C<sub>6</sub> such as CF<sub>3</sub> and CH<sub>2</sub>CH<sub>2</sub>F, lower alkenyl of C<sub>2</sub>-C<sub>6</sub>  
 20 such as CH=CH<sub>2</sub>, halogenated (F, Cl, Br, I) lower alkenyl of C<sub>2</sub>-C<sub>6</sub>  
 such as CH=CHCl, CH=CHBr and CH=CHI, lower alkynyl of C<sub>2</sub>-C<sub>6</sub>  
 such as C≡CH, halogenated (F, Cl, Br, I) lower alkynyl of C<sub>2</sub>-C<sub>6</sub>, lower  
 alkoxy of C<sub>1</sub>-C<sub>6</sub> such as CH<sub>2</sub>OH and CH<sub>2</sub>CH<sub>2</sub>OH, halogenated (F, Cl,  
 Br, I) lower alkoxy of C<sub>1</sub>-C<sub>6</sub>, CO<sub>2</sub>H, CO<sub>2</sub>R', CONH<sub>2</sub>, CONHR',  
 25 CONR'<sub>2</sub>, CH=CHCO<sub>2</sub>H, CH=CHCO<sub>2</sub>R'; and,

R' is an optionally substituted alkyl of C<sub>1</sub>-C<sub>12</sub> (particularly when the alkyl is an  
 amino acid residue), cycloalkyl, optionally substituted alkynyl of C<sub>2</sub>-  
 C<sub>6</sub>, optionally substituted lower alkenyl of C<sub>2</sub>-C<sub>6</sub>, or optionally  
 substituted acyl.

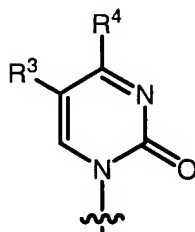
$R^6$  is an optionally substituted alkyl (including lower alkyl), cyano (CN),  $CH_3$ ,  $OCH_3$ ,  $OCH_2CH_3$ , hydroxy methyl ( $CH_2OH$ ), fluoromethyl ( $CH_2F$ ), azido ( $N_3$ ),  $CHCN$ ,  $CH_2N_3$ ,  $CH_2NH_2$ ,  $CH_2NHCH_3$ ,  $CH_2N(CH_3)_2$ , alkyne (optionally substituted), or fluoro;

or its pharmaceutically acceptable salt or prodrug thereof.

10. A (2'R)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside ( $\beta$ -D or  $\beta$ -L) of the formula



wherein Base is



$R^1$  and  $R^7$  are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid (or racemic mixture), a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in*

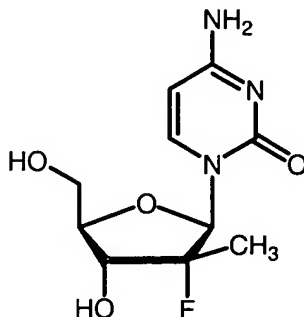
*vivo* is capable of providing a compound wherein R<sup>1</sup> is H or phosphate; R<sup>2</sup> is H or phosphate; R<sup>1</sup> and R<sup>2</sup> or R<sup>7</sup> can also be linked with cyclic phosphate group;

R<sup>3</sup> and R<sup>4</sup> are independently H, halogen including F, Cl, Br, I, OH, OR', SH, SR', NH<sub>2</sub>, NHR', NR'<sub>2</sub>, lower alkyl of C<sub>1</sub>-C<sub>6</sub>, halogenated (F, Cl, Br, I) lower alkyl of C<sub>1</sub>-C<sub>6</sub> such as CF<sub>3</sub> and CH<sub>2</sub>CH<sub>2</sub>F, lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CH<sub>2</sub>, halogenated (F, Cl, Br, I) lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CHCl, CH=CHBr and CH=CHI, lower alkynyl of C<sub>2</sub>-C<sub>6</sub> such as C≡CH, halogenated (F, Cl, Br, I) lower alkynyl of C<sub>2</sub>-C<sub>6</sub>, lower alkoxy of C<sub>1</sub>-C<sub>6</sub> such as CH<sub>2</sub>OH and CH<sub>2</sub>CH<sub>2</sub>OH, halogenated (F, Cl, Br, I) lower alkoxy of C<sub>1</sub>-C<sub>6</sub>, CO<sub>2</sub>H, CO<sub>2</sub>R', CONH<sub>2</sub>, CONHR', CONR'<sub>2</sub>, CH=CHCO<sub>2</sub>H, CH=CHCO<sub>2</sub>R';

R' is an optionally substituted alkyl of C<sub>1</sub>-C<sub>12</sub> (particularly when the alkyl is an amino acid residue), cycloalkyl, optionally substituted alkynyl of C<sub>2</sub>-C<sub>6</sub>, optionally substituted lower alkenyl of C<sub>2</sub>-C<sub>6</sub>, or optionally substituted acyl;

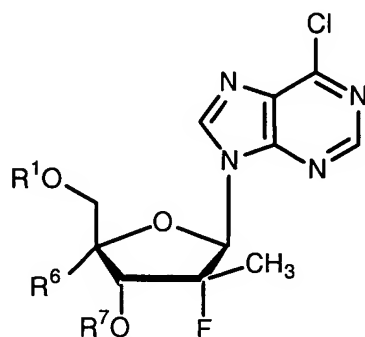
or its pharmaceutically acceptable salt or prodrug thereof.

11. A (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside (β-D) or its pharmaceutically acceptable salt or prodrug thereof of the formula:





12. A (2'R)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside ( $\beta$ -D) or its pharmaceutically acceptable salt or prodrug thereof of the formula:

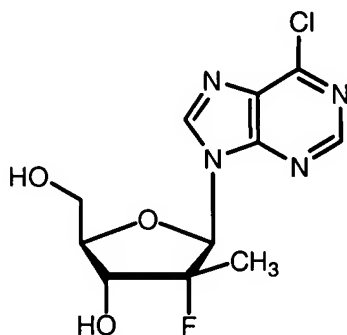


wherein

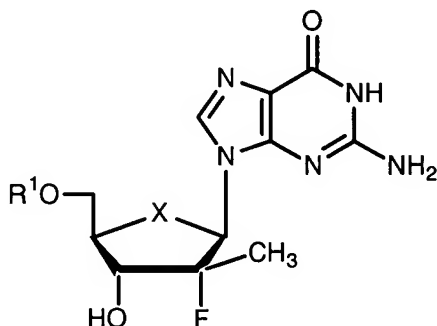
5  $R^1$  and  $R^7$  are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including  
10 methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid, a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein  $R^1$  or  $R^7$  is independently H or  
15 phosphate;  $R^1$  and  $R^7$  can also be linked with cyclic phosphate group; and,

$R^6$  is an optionally substituted alkyl (including lower alkyl), cyano (CN),  $CH_3$ ,  $OCH_3$ ,  $OCH_2CH_3$ , hydroxy methyl ( $CH_2OH$ ), fluoromethyl ( $CH_2F$ ),  
20 azido ( $N_3$ ),  $CHCN$ ,  $CH_2N_3$ ,  $CH_2NH_2$ ,  $CH_2NHCH_3$ ,  $CH_2N(CH_3)_2$ , alkyne (optionally substituted), or fluoro.

13. A (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside ( $\beta$ -D) or its pharmaceutically acceptable salt or prodrug thereof of the formula:



14. A (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside ( $\beta$ -D) or its pharmaceutically acceptable salt or prodrug thereof of the formula:



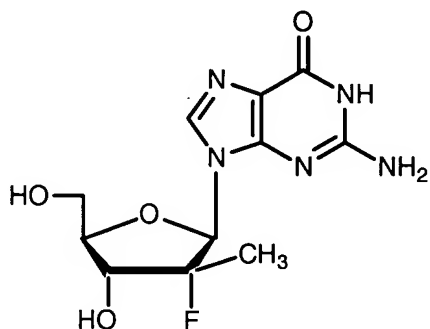
wherein

X is O, S, CH<sub>2</sub>, Se, NH, N-alkyl, CHW (*R*, *S*, or racemic), C(W)<sub>2</sub>, wherein W is F, Cl, Br, or I; and

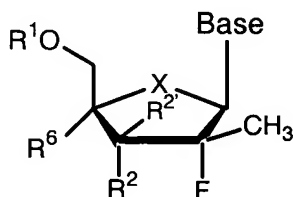
R<sup>1</sup> is H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid, a carbohydrate, a peptide, a cholesterol, or other

pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein R<sup>1</sup> is H or phosphate.

15. A (2'*R*)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside (β-D) or its pharmaceutically acceptable salt or prodrug thereof of the formula:



16. A pharmaceutical composition comprising a (2'*R*)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside (β-D or β-L) of the formula:



wherein

Base is a purine or pyrimidine base;

X is O, S, CH<sub>2</sub>, Se, NH, N-alkyl, CHW (*R*, *S*, or racemic), C(W)<sub>2</sub>, wherein W is F, Cl, Br, or I;

R<sup>1</sup> and R<sup>7</sup> are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives,

sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid (or racemic mixture), a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein R<sup>1</sup> is H or phosphate; R<sup>2</sup> is H or phosphate; R<sup>1</sup> and R<sup>2</sup> or R<sup>7</sup> can also be linked with cyclic phosphate group;

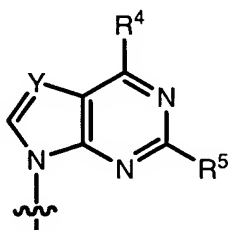
R<sup>2</sup> and R<sup>2'</sup> are independently H, C<sub>1-4</sub> alkyl, C<sub>1-4</sub> alkenyl, C<sub>1-4</sub> alkynyl, vinyl, N<sub>3</sub>, CN, Cl, Br, F, I, NO<sub>2</sub>, C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkynyl), C(O)O(C<sub>1-4</sub> alkenyl), O(C<sub>1-4</sub> acyl), O(C<sub>1-4</sub> alkyl), O(C<sub>1-4</sub> alkenyl), S(C<sub>1-4</sub> acyl), S(C<sub>1-4</sub> alkyl), S(C<sub>1-4</sub> alkynyl), S(C<sub>1-4</sub> alkenyl), SO(C<sub>1-4</sub> acyl), SO(C<sub>1-4</sub> alkyl), SO(C<sub>1-4</sub> alkynyl), SO(C<sub>1-4</sub> alkenyl), SO<sub>2</sub>(C<sub>1-4</sub> acyl), SO<sub>2</sub>(C<sub>1-4</sub> alkyl), SO<sub>2</sub>(C<sub>1-4</sub> alkynyl), SO<sub>2</sub>(C<sub>1-4</sub> alkenyl), O<sub>3</sub>S(C<sub>1-4</sub> acyl), O<sub>3</sub>S(C<sub>1-4</sub> alkyl), O<sub>3</sub>S(C<sub>1-4</sub> alkenyl), NH<sub>2</sub>, NH(C<sub>1-4</sub> alkyl), NH(C<sub>1-4</sub> alkenyl), NH(C<sub>1-4</sub> alkynyl), NH(C<sub>1-4</sub> acyl), N(C<sub>1-4</sub> alkyl)<sub>2</sub>, N(C<sub>1-18</sub> acyl)<sub>2</sub>, wherein alkyl, alkynyl, alkenyl and vinyl are optionally substituted by N<sub>3</sub>, CN, one to three halogen (Cl, Br, F, I), NO<sub>2</sub>, C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkynyl), C(O)O(C<sub>1-4</sub> alkenyl), O(C<sub>1-4</sub> acyl), O(C<sub>1-4</sub> alkyl), O(C<sub>1-4</sub> alkenyl), S(C<sub>1-4</sub> acyl), S(C<sub>1-4</sub> alkyl), S(C<sub>1-4</sub> alkynyl), S(C<sub>1-4</sub> alkenyl), SO(C<sub>1-4</sub> acyl), SO(C<sub>1-4</sub> alkyl), SO(C<sub>1-4</sub> alkynyl), SO(C<sub>1-4</sub> alkenyl), SO<sub>2</sub>(C<sub>1-4</sub> acyl), SO<sub>2</sub>(C<sub>1-4</sub> alkyl), SO<sub>2</sub>(C<sub>1-4</sub> alkynyl), SO<sub>2</sub>(C<sub>1-4</sub> alkenyl), O<sub>3</sub>S(C<sub>1-4</sub> acyl), O<sub>3</sub>S(C<sub>1-4</sub> alkyl), O<sub>3</sub>S(C<sub>1-4</sub> alkenyl), NH<sub>2</sub>, NH(C<sub>1-4</sub> alkyl), NH(C<sub>1-4</sub> alkenyl), NH(C<sub>1-4</sub> alkynyl), NH(C<sub>1-4</sub> acyl), N(C<sub>1-4</sub> alkyl)<sub>2</sub>, N(C<sub>1-4</sub> acyl)<sub>2</sub>, OR<sup>7</sup>; R<sup>2</sup> and R<sup>2'</sup> can be linked together to form a vinyl optionally substituted by one or two of N<sub>3</sub>, CN, Cl, Br, F, I, NO<sub>2</sub>;

R<sup>6</sup> is an optionally substituted alkyl (including lower alkyl), cyano (CN), CH<sub>3</sub>, OCH<sub>3</sub>, OCH<sub>2</sub>CH<sub>3</sub>, hydroxy methyl (CH<sub>2</sub>OH), fluoromethyl (CH<sub>2</sub>F),

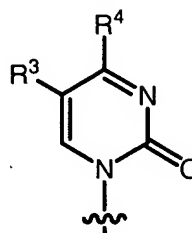
azido ( $N_3$ ),  $CHCN$ ,  $CH_2N_3$ ,  $CH_2NH_2$ ,  $CH_2NHCH_3$ ,  $CH_2N(CH_3)_2$ , alkyne (optionally substituted), or fluoro;

or its pharmaceutically acceptable salt or prodrug thereof, a pharmaceutically acceptable carrier.

- 5            17. The composition of claim 16, wherein Base is selected from the group consisting of:



(a)



(b)

wherein

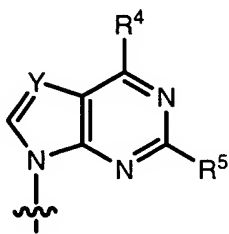
10            Y is N or CH.

15             $R^3$ ,  $R^4$  and  $R^5$  are independently H, halogen including F, Cl, Br, I, OH,  $OR'$ , SH,  $SR'$ ,  $NH_2$ ,  $NHR'$ ,  $NR'_2$ , lower alkyl of  $C_1$ - $C_6$ , halogenated (F, Cl, Br, I) lower alkyl of  $C_1$ - $C_6$  such as  $CF_3$  and  $CH_2CH_2F$ , lower alkenyl of  $C_2$ - $C_6$  such as  $CH=CH_2$ , halogenated (F, Cl, Br, I) lower alkenyl of  $C_2$ - $C_6$  such as  $CH=CHCl$ ,  $CH=CHBr$  and  $CH=CHI$ , lower alkynyl of  $C_2$ - $C_6$  such as  $C\equiv CH$ , halogenated (F, Cl, Br, I) lower alkynyl of  $C_2$ - $C_6$ , lower alkoxy of  $C_1$ - $C_6$  such as  $CH_2OH$  and  $CH_2CH_2OH$ , halogenated (F, Cl, Br, I) lower alkoxy of  $C_1$ - $C_6$ ,  $CO_2H$ ,  $CO_2R'$ ,  $CONH_2$ ,  $CONHR'$ ,  $CONR'_2$ ,  $CH=CHCO_2H$ ,  $CH=CHCO_2R'$ ; and,

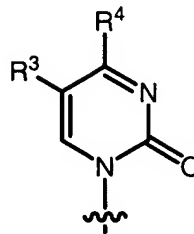
20             $R'$  is an optionally substituted alkyl of  $C_1$ - $C_{12}$  (particularly when the alkyl is an amino acid residue), cycloalkyl, optionally substituted alkynyl of  $C_2$ - $C_6$ , optionally substituted lower alkenyl of  $C_2$ - $C_6$ , or optionally substituted acyl.

18. The composition of claim 16, wherein

Base is selected from the group consisting of (a) or (b):



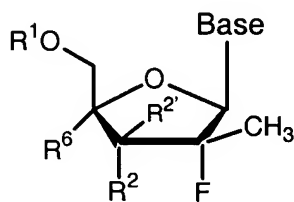
(a)



(b)

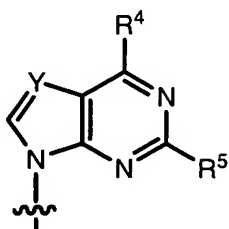
5 and wherein  $R^1$  is H,  $R^2$  is OH,  $R^{2'}$  is H,  $R^3$  is H, and  $R^4$  is  $NH_2$  or OH, and  $R^5$  is  $NH_2$ .

19. A pharmaceutical composition comprising a (2'R)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside ( $\beta$ -D or  $\beta$ -L) of the formula:

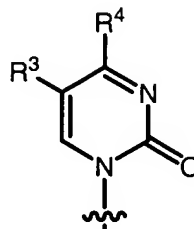


wherein

Base is selected from the group consisting of



(a)



(b)

Y is N or CH;

R<sup>1</sup> and R<sup>7</sup> are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid (or racemic mixture), a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein R<sup>1</sup> is H or phosphate; R<sup>2</sup> is H or phosphate; R<sup>1</sup> and R<sup>2</sup> or R<sup>7</sup> can also be linked with cyclic phosphate group;

R<sup>2</sup> and R<sup>2'</sup> are independently H, C<sub>1-4</sub> alkyl, C<sub>1-4</sub> alkenyl, C<sub>1-4</sub> alkynyl, vinyl, N<sub>3</sub>, CN, Cl, Br, F, I, NO<sub>2</sub>, C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkynyl), C(O)O(C<sub>1-4</sub> alkenyl), O(C<sub>1-4</sub> acyl), O(C<sub>1-4</sub> alkyl), O(C<sub>1-4</sub> alkenyl), S(C<sub>1-4</sub> acyl), S(C<sub>1-4</sub> alkyl), S(C<sub>1-4</sub> alkynyl), S(C<sub>1-4</sub> alkenyl), SO(C<sub>1-4</sub> acyl), SO(C<sub>1-4</sub> alkyl), SO(C<sub>1-4</sub> alkynyl), SO(C<sub>1-4</sub> alkenyl), SO<sub>2</sub>(C<sub>1-4</sub> acyl), SO<sub>2</sub>(C<sub>1-4</sub> alkyl), SO<sub>2</sub>(C<sub>1-4</sub> alkynyl), SO<sub>2</sub>(C<sub>1-4</sub> alkenyl), O<sub>3</sub>S(C<sub>1-4</sub> acyl), O<sub>3</sub>S(C<sub>1-4</sub> alkyl), O<sub>3</sub>S(C<sub>1-4</sub> alkenyl), NH<sub>2</sub>, NH(C<sub>1-4</sub> alkyl), NH(C<sub>1-4</sub> alkenyl), NH(C<sub>1-4</sub> alkynyl), NH(C<sub>1-4</sub> acyl), N(C<sub>1-4</sub> alkyl)<sub>2</sub>, N(C<sub>1-18</sub> acyl)<sub>2</sub>, wherein alkyl, alkynyl, alkenyl and vinyl are optionally substituted by N<sub>3</sub>, CN, one to three halogen (Cl, Br, F, I), NO<sub>2</sub>, C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkynyl), C(O)O(C<sub>1-4</sub> alkenyl), O(C<sub>1-4</sub> acyl), O(C<sub>1-4</sub> alkyl), O(C<sub>1-4</sub> alkenyl), S(C<sub>1-4</sub> acyl), S(C<sub>1-4</sub> alkyl), S(C<sub>1-4</sub> alkynyl), S(C<sub>1-4</sub> alkenyl), SO(C<sub>1-4</sub> acyl), SO(C<sub>1-4</sub> alkyl), SO(C<sub>1-4</sub> alkynyl), SO(C<sub>1-4</sub> alkenyl), SO<sub>2</sub>(C<sub>1-4</sub> acyl), SO<sub>2</sub>(C<sub>1-4</sub> alkyl), SO<sub>2</sub>(C<sub>1-4</sub> alkynyl), SO<sub>2</sub>(C<sub>1-4</sub> alkenyl), O<sub>3</sub>S(C<sub>1-4</sub> acyl), O<sub>3</sub>S(C<sub>1-4</sub> alkyl), O<sub>3</sub>S(C<sub>1-4</sub> alkenyl), NH<sub>2</sub>, NH(C<sub>1-4</sub> alkyl), NH(C<sub>1-4</sub>

alkenyl), NH(C<sub>1-4</sub> alkynyl), NH(C<sub>1-4</sub> acyl), N(C<sub>1-4</sub> alkyl)<sub>2</sub>, N(C<sub>1-4</sub> acyl)<sub>2</sub>, OR<sup>7</sup>; R<sup>2</sup> and R<sup>2'</sup> can be linked together to form a vinyl optionally substituted by one or two of N<sub>3</sub>, CN, Cl, Br, F, I, NO<sub>2</sub>;

5 R<sup>3</sup>, R<sup>4</sup> and R<sup>5</sup> are independently H, halogen including F, Cl, Br, I, OH, OR', SH, SR', NH<sub>2</sub>, NHR', NR'<sub>2</sub>, lower alkyl of C<sub>1</sub>-C<sub>6</sub>, halogenated (F, Cl, Br, I) lower alkyl of C<sub>1</sub>-C<sub>6</sub> such as CF<sub>3</sub> and CH<sub>2</sub>CH<sub>2</sub>F, lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CH<sub>2</sub>, halogenated (F, Cl, Br, I) lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CHCl, CH=CHBr and CH=CHI, lower alkynyl of C<sub>2</sub>-C<sub>6</sub> such as C≡CH, halogenated (F, Cl, Br, I) lower alkynyl of C<sub>2</sub>-C<sub>6</sub>,  
10 lower alkoxy of C<sub>1</sub>-C<sub>6</sub> such as CH<sub>2</sub>OH and CH<sub>2</sub>CH<sub>2</sub>OH, halogenated (F, Cl, Br, I) lower alkoxy of C<sub>1</sub>-C<sub>6</sub>, CO<sub>2</sub>H, CO<sub>2</sub>R', CONH<sub>2</sub>, CONHR', CONR'<sub>2</sub>, CH=CHCO<sub>2</sub>H, CH=CHCO<sub>2</sub>R';

15 R' is an optionally substituted alkyl of C<sub>1</sub>-C<sub>12</sub> (particularly when the alkyl is an amino acid residue), cycloalkyl, optionally substituted alkynyl of C<sub>2</sub>-C<sub>6</sub>, optionally substituted lower alkenyl of C<sub>2</sub>-C<sub>6</sub>, or optionally substituted acyl;

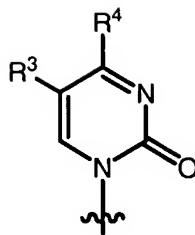
20 R<sup>6</sup> is an optionally substituted alkyl (including lower alkyl), cyano (CN), CH<sub>3</sub>, OCH<sub>3</sub>, OCH<sub>2</sub>CH<sub>3</sub>, hydroxy methyl (CH<sub>2</sub>OH), fluoromethyl (CH<sub>2</sub>F), azido (N<sub>3</sub>), CHCN, CH<sub>2</sub>N<sub>3</sub>, CH<sub>2</sub>NH<sub>2</sub>, CH<sub>2</sub>NHCH<sub>3</sub>, CH<sub>2</sub>N(CH<sub>3</sub>)<sub>2</sub>, alkyne (optionally substituted), or fluoro;

or its pharmaceutically acceptable salt or prodrug thereof in a pharmaceutically acceptable carrier.

20. The composition of claim 19, wherein

25 Base is

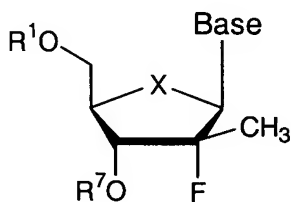




and R<sup>1</sup> is H, R<sup>2</sup> is OH, R<sup>2'</sup> is H, R<sup>3</sup> is H, R<sup>4</sup> is NH<sub>2</sub> or OH,  
and R<sup>6</sup> is H.

5

21. A pharmaceutical composition comprising a (2'*R*)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside (β-D or β-L) or its pharmaceutically acceptable salt or prodrug thereof, in a pharmaceutically acceptable carrier, of the structure:



10

wherein Base is a purine or pyrimidine base;

X is O, S, CH<sub>2</sub>, Se, NH, N-alkyl, CHW (*R*, *S*, or racemic), C(W)<sub>2</sub>, wherein W is F, Cl, Br, or I; and,

15

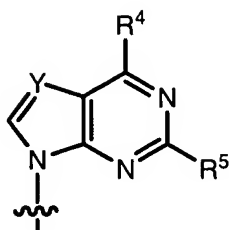
R<sup>1</sup> and R<sup>7</sup> are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid, a carbohydrate, a peptide, a cholesterol, or other pharmaceutically

20

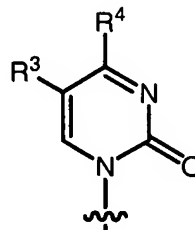
acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein  $R^1$  or  $R^7$  is independently H or phosphate;  $R^1$  and  $R^7$  can also be linked with cyclic phosphate group.

5                    22. The composition of claim 21, wherein

Base is selected from the group consisting of:



(a)



(b)

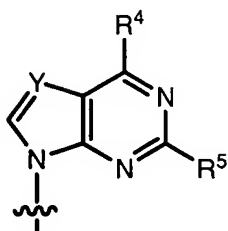
Y is N or CH;

10                     $R^3$ ,  $R^4$  and  $R^5$  are independently H, halogen including F, Cl, Br, I, OH, OR', SH, SR', NH<sub>2</sub>, NHR', NR'<sub>2</sub>, lower alkyl of C<sub>1</sub>-C<sub>6</sub>, halogenated (F, Cl, Br, I) lower alkyl of C<sub>1</sub>-C<sub>6</sub> such as CF<sub>3</sub> and CH<sub>2</sub>CH<sub>2</sub>F, lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CH<sub>2</sub>, halogenated (F, Cl, Br, I) lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CHCl, CH=CHBr and CH=CHI, lower alkynyl of C<sub>2</sub>-C<sub>6</sub> such as C≡CH, halogenated (F, Cl, Br, I) lower alkynyl of C<sub>2</sub>-C<sub>6</sub>, lower alkoxy of C<sub>1</sub>-C<sub>6</sub> such as CH<sub>2</sub>OH and CH<sub>2</sub>CH<sub>2</sub>OH, halogenated (F, Cl, Br, I) lower alkoxy of C<sub>1</sub>-C<sub>6</sub>, CO<sub>2</sub>H, CO<sub>2</sub>R', CONH<sub>2</sub>, CONHR', CONR'<sub>2</sub>, CH=CHCO<sub>2</sub>H, CH=CHCO<sub>2</sub>R'; and,

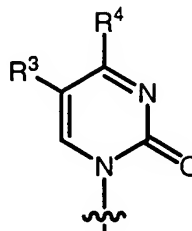
20                    R' is an optionally substituted alkyl of C<sub>1</sub>-C<sub>12</sub> (particularly when the alkyl is an amino acid residue), cycloalkyl, optionally substituted alkynyl of C<sub>2</sub>-C<sub>6</sub>, optionally substituted lower alkenyl of C<sub>2</sub>-C<sub>6</sub>, or optionally substituted acyl.

23. The composition of claim 21, wherein

Base is selected from the group consisting of (a) or (b):



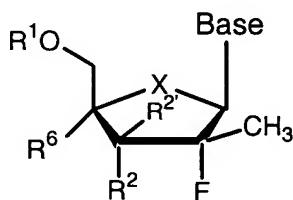
(a)



(b)

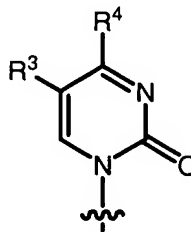
5 and wherein R<sup>1</sup> and R<sup>7</sup> are H, R<sup>3</sup> is H, and R<sup>4</sup> is NH<sub>2</sub> or OH, and R<sup>5</sup> is NH<sub>2</sub>.

24. A pharmaceutical composition comprising a (2'R)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside (β-D or β-L) of the formula:



10 wherein

Base is



X is O, S, CH<sub>2</sub>, Se, NH, N-alkyl, CHW (R, S, or racemic), C(W)<sub>2</sub>, wherein W is F, Cl, Br, or I;

15 R<sup>1</sup> and R<sup>7</sup> are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-

phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid (or racemic mixture), a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein R<sup>1</sup> is H or phosphate; R<sup>2</sup> is H or phosphate; R<sup>1</sup> and R<sup>2</sup> or R<sup>7</sup> can also be linked with cyclic phosphate group;

R<sup>2</sup> and R<sup>2'</sup> are independently H, C<sub>1-4</sub> alkyl, C<sub>1-4</sub> alkenyl, C<sub>1-4</sub> alkynyl, vinyl, N<sub>3</sub>, CN, Cl, Br, F, I, NO<sub>2</sub>, C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkynyl), C(O)O(C<sub>1-4</sub> alkenyl), O(C<sub>1-4</sub> acyl), O(C<sub>1-4</sub> alkyl), O(C<sub>1-4</sub> alkenyl), S(C<sub>1-4</sub> acyl), S(C<sub>1-4</sub> alkyl), S(C<sub>1-4</sub> alkynyl), S(C<sub>1-4</sub> alkenyl), SO(C<sub>1-4</sub> acyl), SO(C<sub>1-4</sub> alkyl), SO(C<sub>1-4</sub> alkynyl), SO(C<sub>1-4</sub> alkenyl), SO<sub>2</sub>(C<sub>1-4</sub> acyl), SO<sub>2</sub>(C<sub>1-4</sub> alkyl), SO<sub>2</sub>(C<sub>1-4</sub> alkynyl), SO<sub>2</sub>(C<sub>1-4</sub> alkenyl), O<sub>3</sub>S(C<sub>1-4</sub> acyl), O<sub>3</sub>S(C<sub>1-4</sub> alkyl), O<sub>3</sub>S(C<sub>1-4</sub> alkenyl), NH<sub>2</sub>, NH(C<sub>1-4</sub> alkyl), NH(C<sub>1-4</sub> alkenyl), NH(C<sub>1-4</sub> alkynyl), NH(C<sub>1-4</sub> acyl), N(C<sub>1-4</sub> alkyl)<sub>2</sub>, N(C<sub>1-18</sub> acyl)<sub>2</sub>, wherein alkyl, alkynyl, alkenyl and vinyl are optionally substituted by N<sub>3</sub>, CN, one to three halogen (Cl, Br, F, I), NO<sub>2</sub>, C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkynyl), C(O)O(C<sub>1-4</sub> alkenyl), O(C<sub>1-4</sub> acyl), O(C<sub>1-4</sub> alkyl), O(C<sub>1-4</sub> alkenyl), S(C<sub>1-4</sub> acyl), S(C<sub>1-4</sub> alkyl), S(C<sub>1-4</sub> alkynyl), S(C<sub>1-4</sub> alkenyl), SO(C<sub>1-4</sub> acyl), SO(C<sub>1-4</sub> alkyl), SO(C<sub>1-4</sub> alkynyl), SO(C<sub>1-4</sub> alkenyl), SO<sub>2</sub>(C<sub>1-4</sub> acyl), SO<sub>2</sub>(C<sub>1-4</sub> alkyl), SO<sub>2</sub>(C<sub>1-4</sub> alkynyl), SO<sub>2</sub>(C<sub>1-4</sub> alkenyl), O<sub>3</sub>S(C<sub>1-4</sub> acyl), O<sub>3</sub>S(C<sub>1-4</sub> alkyl), O<sub>3</sub>S(C<sub>1-4</sub> alkenyl), NH<sub>2</sub>, NH(C<sub>1-4</sub> alkyl), NH(C<sub>1-4</sub> alkenyl), NH(C<sub>1-4</sub> alkynyl), NH(C<sub>1-4</sub> acyl), N(C<sub>1-4</sub> alkyl)<sub>2</sub>, N(C<sub>1-4</sub> acyl)<sub>2</sub>, OR<sup>7</sup>; R<sup>2</sup> and R<sup>2'</sup> can be linked together to form a vinyl optionally substituted by one or two of N<sub>3</sub>, CN, Cl, Br, F, I, NO<sub>2</sub>;

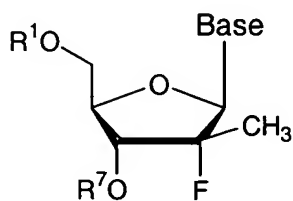
$R^3$  and  $R^4$  are independently H, halogen including F, Cl, Br, I, OH, OR', SH, SR', NH<sub>2</sub>, NHR', NR'<sub>2</sub>, lower alkyl of C<sub>1</sub>-C<sub>6</sub>, halogenated (F, Cl, Br, I) lower alkyl of C<sub>1</sub>-C<sub>6</sub> such as CF<sub>3</sub> and CH<sub>2</sub>CH<sub>2</sub>F, lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CH<sub>2</sub>, halogenated (F, Cl, Br, I) lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CHCl, CH=CHBr and CH=CHI, lower alkynyl of C<sub>2</sub>-C<sub>6</sub> such as C≡CH, halogenated (F, Cl, Br, I) lower alkynyl of C<sub>2</sub>-C<sub>6</sub>, lower alkoxy of C<sub>1</sub>-C<sub>6</sub> such as CH<sub>2</sub>OH and CH<sub>2</sub>CH<sub>2</sub>OH, halogenated (F, Cl, Br, I) lower alkoxy of C<sub>1</sub>-C<sub>6</sub>, CO<sub>2</sub>H, CO<sub>2</sub>R', CONH<sub>2</sub>, CONHR', CONR'<sub>2</sub>, CH=CHCO<sub>2</sub>H, CH=CHCO<sub>2</sub>R';

$R'$  is an optionally substituted alkyl of C<sub>1</sub>-C<sub>12</sub> (particularly when the alkyl is an amino acid residue), cycloalkyl, optionally substituted alkynyl of C<sub>2</sub>-C<sub>6</sub>, optionally substituted lower alkenyl of C<sub>2</sub>-C<sub>6</sub>, or optionally substituted acyl; and

$R^6$  is an optionally substituted alkyl (including lower alkyl), cyano (CN), CH<sub>3</sub>, OCH<sub>3</sub>, OCH<sub>2</sub>CH<sub>3</sub>, hydroxy methyl (CH<sub>2</sub>OH), fluoromethyl (CH<sub>2</sub>F), azido (N<sub>3</sub>), CHCN, CH<sub>2</sub>N<sub>3</sub>, CH<sub>2</sub>NH<sub>2</sub>, CH<sub>2</sub>NHCH<sub>3</sub>, CH<sub>2</sub>N(CH<sub>3</sub>)<sub>2</sub>, alkyne (optionally substituted), or fluoro;

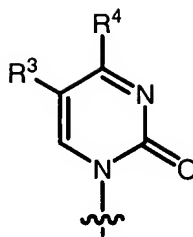
or its pharmaceutically acceptable salt or prodrug thereof and a pharmaceutically acceptable carrier.

25. A pharmaceutical composition comprising a (2'*R*)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside (β-D or β-L) of the formula:



wherein

Base is



$R^1$  and  $R^7$  are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid, a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein  $R^1$  or  $R^7$  is independently H or phosphate;  $R^1$  and  $R^7$  can also be linked with cyclic phosphate group;

$R^3$  and  $R^4$  are independently H, halogen including F, Cl, Br, I, OH, OR', SH, SR', NH<sub>2</sub>, NHR', NR'<sub>2</sub>, lower alkyl of C<sub>1</sub>-C<sub>6</sub>, halogenated (F, Cl, Br, I) lower alkyl of C<sub>1</sub>-C<sub>6</sub> such as CF<sub>3</sub> and CH<sub>2</sub>CH<sub>2</sub>F, lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CH<sub>2</sub>, halogenated (F, Cl, Br, I) lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CHCl, CH=CHBr and CH=CHI, lower alkynyl of C<sub>2</sub>-C<sub>6</sub> such as C≡CH, halogenated (F, Cl, Br, I) lower alkynyl of C<sub>2</sub>-C<sub>6</sub>, lower alkoxy of C<sub>1</sub>-C<sub>6</sub> such as CH<sub>2</sub>OH and CH<sub>2</sub>CH<sub>2</sub>OH, halogenated (F, Cl, Br, I) lower alkoxy of C<sub>1</sub>-C<sub>6</sub>, CO<sub>2</sub>H, CO<sub>2</sub>R', CONH<sub>2</sub>, CONHR', CONR'<sub>2</sub>, CH=CHCO<sub>2</sub>H, CH=CHCO<sub>2</sub>R';

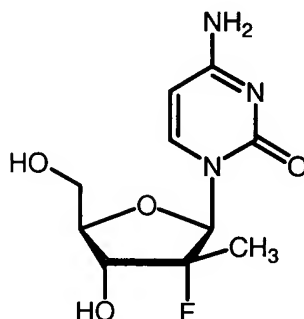
$R'$  is an optionally substituted alkyl of C<sub>1</sub>-C<sub>12</sub> (particularly when the alkyl is an amino acid residue), cycloalkyl, optionally substituted alkynyl of C<sub>2</sub>-C<sub>6</sub>,

optionally substituted lower alkenyl of C<sub>2</sub>-C<sub>6</sub>, or optionally substituted acyl;

or its pharmaceutically acceptable salt or prodrug thereof, in a pharmaceutically acceptable carrier.

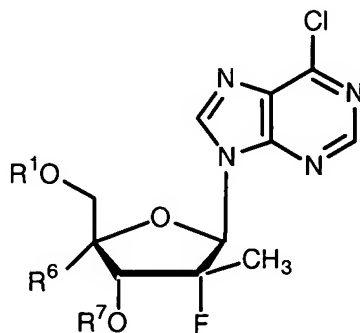
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26. A pharmaceutical composition comprising a (2'*R*)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside (β-D) or its pharmaceutically acceptable salt or prodrug thereof, in a pharmaceutically acceptable carrier of the formula:



10

27. A pharmaceutical composition comprising a (2'*R*)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside (β-D or β-L), or its pharmaceutically acceptable salt or prodrug thereof, in a pharmaceutically acceptable carrier, of the formula:



15

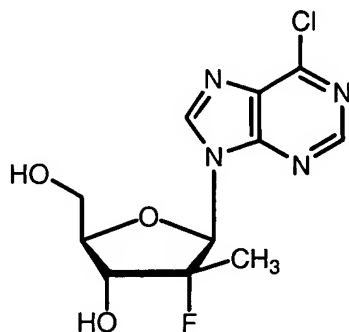
wherein

R<sup>1</sup> and R<sup>7</sup> are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-

phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid, a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein R<sup>1</sup> or R<sup>7</sup> is independently H or phosphate; R<sup>1</sup> and R<sup>7</sup> can also be linked with cyclic phosphate group; and,

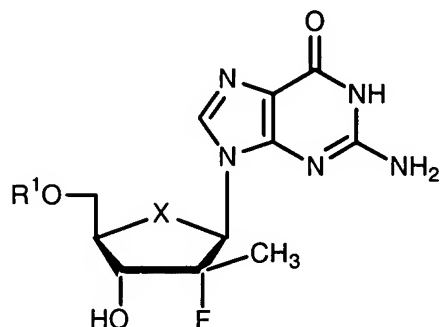
R<sup>6</sup> is an optionally substituted alkyl (including lower alkyl), cyano (CN), CH<sub>3</sub>, OCH<sub>3</sub>, OCH<sub>2</sub>CH<sub>3</sub>, hydroxy methyl (CH<sub>2</sub>OH), fluoromethyl (CH<sub>2</sub>F), azido (N<sub>3</sub>), CHCN, CH<sub>2</sub>N<sub>3</sub>, CH<sub>2</sub>NH<sub>2</sub>, CH<sub>2</sub>NHCH<sub>3</sub>, CH<sub>2</sub>N(CH<sub>3</sub>)<sub>2</sub>, alkyne (optionally substituted), or fluoro.

28. A pharmaceutical composition comprising a (2'*R*)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside (β-D) or its pharmaceutically acceptable salt or prodrug thereof, in a pharmaceutically acceptable carrier, of the formula:





29. A pharmaceutical composition comprising a (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside ( $\beta$ -D) or its pharmaceutically acceptable salt or prodrug thereof in a pharmaceutically acceptable carrier of the formula:



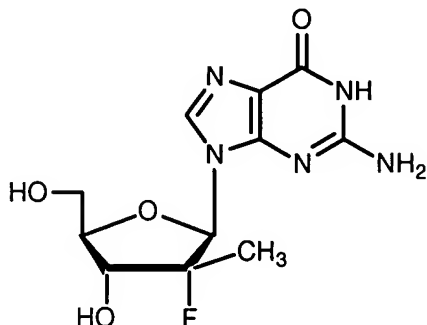
5                    wherein

X is O, S, CH<sub>2</sub>, Se, NH, N-alkyl, CHW (*R*, *S*, or racemic), C(W)<sub>2</sub>, wherein W is F, Cl, Br, or I; and

10                    R<sup>1</sup> is H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid, a carbohydrate, a peptide, a cholesterol, or other  
15                    pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein R<sup>1</sup> is H or phosphate;

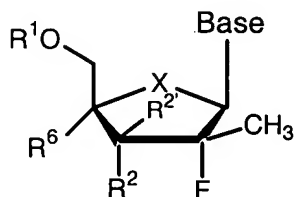
20

30. A pharmaceutical composition comprising a (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside ( $\beta$ -D) or its pharmaceutically acceptable salt or prodrug thereof, in a pharmaceutically acceptable carrier, of the structure:



5

31. A method for the treatment or prophylaxis of hepatitis C infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside ( $\beta$ -D or  $\beta$ -L) of the formula:



10

wherein

Base is a purine or pyrimidine base;

X is O, S, CH<sub>2</sub>, Se, NH, N-alkyl, CHW (*R*, *S*, or racemic), C(W)<sub>2</sub>, wherein W is F, Cl, Br, or I;

15  $R^1$  and  $R^7$  are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives,

sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid (or racemic mixture), a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein R<sup>1</sup> is H or phosphate; R<sup>2</sup> is H or phosphate; R<sup>1</sup> and R<sup>2</sup> or R<sup>7</sup> can also be linked with cyclic phosphate group;

R<sup>2</sup> and R<sup>2'</sup> are independently H, C<sub>1-4</sub> alkyl, C<sub>1-4</sub> alkenyl, C<sub>1-4</sub> alkynyl, vinyl, N<sub>3</sub>, CN, Cl, Br, F, I, NO<sub>2</sub>, C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkynyl), C(O)O(C<sub>1-4</sub> alkenyl), O(C<sub>1-4</sub> acyl), O(C<sub>1-4</sub> alkyl), O(C<sub>1-4</sub> alkenyl), S(C<sub>1-4</sub> acyl), S(C<sub>1-4</sub> alkyl), S(C<sub>1-4</sub> alkynyl), S(C<sub>1-4</sub> alkenyl), SO(C<sub>1-4</sub> acyl), SO(C<sub>1-4</sub> alkyl), SO(C<sub>1-4</sub> alkynyl), SO(C<sub>1-4</sub> alkenyl), SO<sub>2</sub>(C<sub>1-4</sub> acyl), SO<sub>2</sub>(C<sub>1-4</sub> alkyl), SO<sub>2</sub>(C<sub>1-4</sub> alkynyl), SO<sub>2</sub>(C<sub>1-4</sub> alkenyl), O<sub>3</sub>S(C<sub>1-4</sub> acyl), O<sub>3</sub>S(C<sub>1-4</sub> alkyl), O<sub>3</sub>S(C<sub>1-4</sub> alkenyl), NH<sub>2</sub>, NH(C<sub>1-4</sub> alkyl), NH(C<sub>1-4</sub> alkenyl), NH(C<sub>1-4</sub> alkynyl), NH(C<sub>1-4</sub> acyl), N(C<sub>1-4</sub> alkyl)<sub>2</sub>, N(C<sub>1-18</sub> acyl)<sub>2</sub>, wherein alkyl, alkynyl, alkenyl and vinyl are optionally substituted by N<sub>3</sub>, CN, one to three halogen (Cl, Br, F, I), NO<sub>2</sub>, C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkynyl), C(O)O(C<sub>1-4</sub> alkenyl), O(C<sub>1-4</sub> acyl), O(C<sub>1-4</sub> alkyl), O(C<sub>1-4</sub> alkenyl), S(C<sub>1-4</sub> acyl), S(C<sub>1-4</sub> alkyl), S(C<sub>1-4</sub> alkynyl), S(C<sub>1-4</sub> alkenyl), SO(C<sub>1-4</sub> acyl), SO(C<sub>1-4</sub> alkyl), SO(C<sub>1-4</sub> alkynyl), SO(C<sub>1-4</sub> alkenyl), SO<sub>2</sub>(C<sub>1-4</sub> acyl), SO<sub>2</sub>(C<sub>1-4</sub> alkyl), SO<sub>2</sub>(C<sub>1-4</sub> alkynyl), SO<sub>2</sub>(C<sub>1-4</sub> alkenyl), O<sub>3</sub>S(C<sub>1-4</sub> acyl), O<sub>3</sub>S(C<sub>1-4</sub> alkyl), O<sub>3</sub>S(C<sub>1-4</sub> alkenyl), NH<sub>2</sub>, NH(C<sub>1-4</sub> alkyl), NH(C<sub>1-4</sub> alkenyl), NH(C<sub>1-4</sub> alkynyl), NH(C<sub>1-4</sub> acyl), N(C<sub>1-4</sub> alkyl)<sub>2</sub>, N(C<sub>1-4</sub> acyl)<sub>2</sub>, OR<sup>7</sup>; R<sup>2</sup> and R<sup>2'</sup> can be linked together to form a vinyl optionally substituted by one or two of N<sub>3</sub>, CN, Cl, Br, F, I, NO<sub>2</sub>;

R<sup>6</sup> is an optionally substituted alkyl (including lower alkyl), cyano (CN), CH<sub>3</sub>, OCH<sub>3</sub>, OCH<sub>2</sub>CH<sub>3</sub>, hydroxy methyl (CH<sub>2</sub>OH), fluoromethyl (CH<sub>2</sub>F),

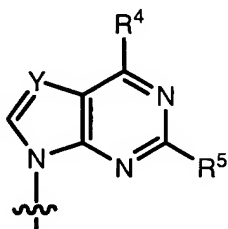
azido ( $N_3$ ),  $CHCN$ ,  $CH_2N_3$ ,  $CH_2NH_2$ ,  $CH_2NHCH_3$ ,  $CH_2N(CH_3)_2$ ,  
alkyne (optionally substituted), or fluoro;

or its pharmaceutically acceptable salt or prodrug thereof, optionally in a pharmaceutically acceptable carrier.

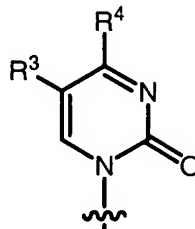
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32. The method of claim 31,

wherein Base is selected from the group consisting of:



(a)



(b)

10

Y is N or CH.

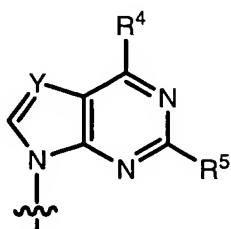
$R^3$ ,  $R^4$  and  $R^5$  are independently H, halogen including F, Cl, Br, I, OH,  $OR'$ ,  
SH,  $SR'$ ,  $NH_2$ ,  $NHR'$ ,  $NR'_2$ , lower alkyl of  $C_1$ - $C_6$ , halogenated (F, Cl,  
Br, I) lower alkyl of  $C_1$ - $C_6$  such as  $CF_3$  and  $CH_2CH_2F$ , lower alkenyl of  
15  $C_2$ - $C_6$  such as  $CH=CH_2$ , halogenated (F, Cl, Br, I) lower alkenyl of  $C_2$ -  
 $C_6$  such as  $CH=CHCl$ ,  $CH=CHBr$  and  $CH=CHI$ , lower alkynyl of  $C_2$ -  
 $C_6$  such as  $C\equiv CH$ , halogenated (F, Cl, Br, I) lower alkynyl of  $C_2$ - $C_6$ ,  
lower alkoxy of  $C_1$ - $C_6$  such as  $CH_2OH$  and  $CH_2CH_2OH$ , halogenated  
(F, Cl, Br, I) lower alkoxy of  $C_1$ - $C_6$ ,  $CO_2H$ ,  $CO_2R'$ ,  $CONH_2$ ,  $CONHR'$ ,  
20  $CONR'_2$ ,  $CH=CHCO_2H$ ,  $CH=CHCO_2R'$ ; and,

$R'$  is an optionally substituted alkyl of  $C_1$ - $C_{12}$  (particularly when the alkyl is an  
amino acid residue), cycloalkyl, optionally substituted alkynyl of  $C_2$ -

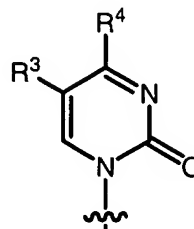
C<sub>6</sub>, optionally substituted lower alkenyl of C<sub>2</sub>-C<sub>6</sub>, or optionally substituted acyl.

33. The method of claim 31, wherein

5 Base is selected from the group consisting of (a) or (b):



(a)

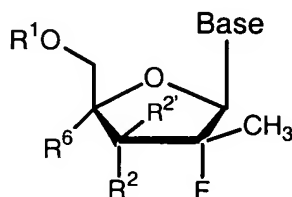


(b)

and wherein R<sup>1</sup> is H, R<sup>2</sup> is OH, R<sup>2'</sup> is H, R<sup>3</sup> is H, and R<sup>4</sup> is NH<sub>2</sub> or OH, and R<sup>5</sup> is NH<sub>2</sub>.

10

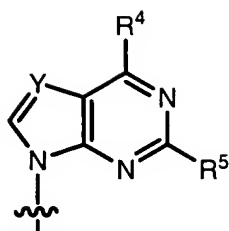
34. A method for the treatment or prophylaxis of hepatitis C infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside (β-D or β-L) of the formula:



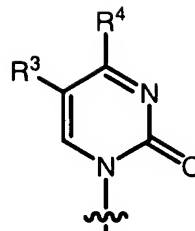
15

wherein

Base is selected from the group consisting of



(a)



(b)

Y is N or CH;

R<sup>1</sup> and R<sup>7</sup> are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid (or racemic mixture), a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein R<sup>1</sup> is H or phosphate; R<sup>2</sup> is H or phosphate; R<sup>1</sup> and R<sup>2</sup> or R<sup>7</sup> can also be linked with cyclic phosphate group;

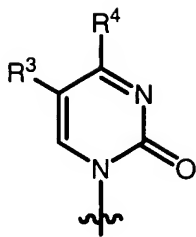
R<sup>2</sup> and R<sup>2'</sup> are independently H, C<sub>1-4</sub> alkyl, C<sub>1-4</sub> alkenyl, C<sub>1-4</sub> alkynyl, vinyl, N<sub>3</sub>, CN, Cl, Br, F, I, NO<sub>2</sub>, C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkynyl), C(O)O(C<sub>1-4</sub> alkenyl), O(C<sub>1-4</sub> acyl), O(C<sub>1-4</sub> alkyl), O(C<sub>1-4</sub> alkenyl), S(C<sub>1-4</sub> acyl), S(C<sub>1-4</sub> alkyl), S(C<sub>1-4</sub> alkynyl), S(C<sub>1-4</sub> alkenyl), SO(C<sub>1-4</sub> acyl), SO(C<sub>1-4</sub> alkyl), SO(C<sub>1-4</sub> alkynyl), SO(C<sub>1-4</sub> alkenyl), SO<sub>2</sub>(C<sub>1-4</sub> acyl), SO<sub>2</sub>(C<sub>1-4</sub> alkyl), SO<sub>2</sub>(C<sub>1-4</sub> alkynyl), SO<sub>2</sub>(C<sub>1-4</sub> alkenyl), O<sub>3</sub>S(C<sub>1-4</sub> acyl), O<sub>3</sub>S(C<sub>1-4</sub> alkyl), O<sub>3</sub>S(C<sub>1-4</sub> alkenyl), NH<sub>2</sub>, NH(C<sub>1-4</sub> alkyl), NH(C<sub>1-4</sub> alkenyl), NH(C<sub>1-4</sub> alkynyl), NH(C<sub>1-4</sub> acyl), N(C<sub>1-4</sub> alkyl)<sub>2</sub>, N(C<sub>1-18</sub> acyl)<sub>2</sub>, wherein alkyl, alkynyl, alkenyl and vinyl are optionally substituted by N<sub>3</sub>, CN, one to three halogen (Cl, Br, F, I),

- 5 NO<sub>2</sub>, C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkynyl), C(O)O(C<sub>1-4</sub> alkenyl), O(C<sub>1-4</sub> acyl), O(C<sub>1-4</sub> alkyl), O(C<sub>1-4</sub> alkenyl), S(C<sub>1-4</sub> acyl), S(C<sub>1-4</sub> alkyl), S(C<sub>1-4</sub> alkynyl), S(C<sub>1-4</sub> alkenyl), SO(C<sub>1-4</sub> acyl), SO(C<sub>1-4</sub> alkyl), SO(C<sub>1-4</sub> alkynyl), SO(C<sub>1-4</sub> alkenyl), SO<sub>2</sub>(C<sub>1-4</sub> acyl), SO<sub>2</sub>(C<sub>1-4</sub> alkyl), SO<sub>2</sub>(C<sub>1-4</sub> alkynyl), SO<sub>2</sub>(C<sub>1-4</sub> alkenyl), O<sub>3</sub>S(C<sub>1-4</sub> acyl), O<sub>3</sub>S(C<sub>1-4</sub> alkyl), O<sub>3</sub>S(C<sub>1-4</sub> alkenyl), NH<sub>2</sub>, NH(C<sub>1-4</sub> alkyl), NH(C<sub>1-4</sub> alkenyl), NH(C<sub>1-4</sub> alkynyl), NH(C<sub>1-4</sub> acyl), N(C<sub>1-4</sub> alkyl)<sub>2</sub>, N(C<sub>1-4</sub> acyl)<sub>2</sub>, OR<sup>7</sup>; R<sup>2</sup> and R<sup>2'</sup> can be linked together to form a vinyl optionally substituted by one or two of N<sub>3</sub>, CN, Cl, Br, F, I, NO<sub>2</sub>;
- 10 R<sup>3</sup>, R<sup>4</sup> and R<sup>5</sup> are independently H, halogen including F, Cl, Br, I, OH, OR', SH, SR', NH<sub>2</sub>, NHR', NR'<sub>2</sub>, lower alkyl of C<sub>1</sub>-C<sub>6</sub>, halogenated (F, Cl, Br, I) lower alkyl of C<sub>1</sub>-C<sub>6</sub> such as CF<sub>3</sub> and CH<sub>2</sub>CH<sub>2</sub>F, lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CH<sub>2</sub>, halogenated (F, Cl, Br, I) lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CHCl, CH=CHBr and CH=CHI, lower alkynyl of C<sub>2</sub>-C<sub>6</sub> such as C≡CH, halogenated (F, Cl, Br, I) lower alkynyl of C<sub>2</sub>-C<sub>6</sub>, lower alkoxy of C<sub>1</sub>-C<sub>6</sub> such as CH<sub>2</sub>OH and CH<sub>2</sub>CH<sub>2</sub>OH, halogenated (F, Cl, Br, I) lower alkoxy of C<sub>1</sub>-C<sub>6</sub>, CO<sub>2</sub>H, CO<sub>2</sub>R', CONH<sub>2</sub>, CONHR', CONR'<sub>2</sub>, CH=CHCO<sub>2</sub>H, CH=CHCO<sub>2</sub>R';
- 15
- 20 R' is an optionally substituted alkyl of C<sub>1</sub>-C<sub>12</sub> (particularly when the alkyl is an amino acid residue), cycloalkyl, optionally substituted alkynyl of C<sub>2</sub>-C<sub>6</sub>, optionally substituted lower alkenyl of C<sub>2</sub>-C<sub>6</sub>, or optionally substituted acyl;
- 25 R<sup>6</sup> is an optionally substituted alkyl (including lower alkyl), cyano (CN), CH<sub>3</sub>, OCH<sub>3</sub>, OCH<sub>2</sub>CH<sub>3</sub>, hydroxy methyl (CH<sub>2</sub>OH), fluoromethyl (CH<sub>2</sub>F), azido (N<sub>3</sub>), CHCN, CH<sub>2</sub>N<sub>3</sub>, CH<sub>2</sub>NH<sub>2</sub>, CH<sub>2</sub>NHCH<sub>3</sub>, CH<sub>2</sub>N(CH<sub>3</sub>)<sub>2</sub>, alkyne (optionally substituted), or fluoro;

or its pharmaceutically acceptable salt or prodrug thereof, optionally in a pharmaceutically acceptable carrier.

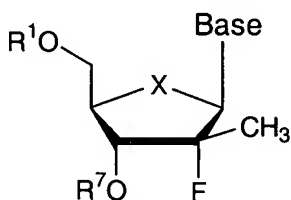
35. The method of claim 34, wherein

Base is



5 and R<sup>1</sup> is H, R<sup>2</sup> is OH, R<sup>2'</sup> is H, R<sup>3</sup> is H, R<sup>4</sup> is NH<sub>2</sub> or OH,  
and R<sup>6</sup> is H.

36. A method for the treatment or prophylaxis of hepatitis C infection comprising  
administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-C-  
10 methyl nucleoside (β-D or β-L) or its pharmaceutically acceptable salt or prodrug thereof of  
the structure:



wherein Base is a purine or pyrimidine base;

15 X is O, S, CH<sub>2</sub>, Se, NH, N-alkyl, CHW (*R*, *S*, or racemic), C(W)<sub>2</sub>, wherein W  
is F, Cl, Br, or I; and,

R<sup>1</sup> and R<sup>7</sup> are independently H, phosphate, including monophosphate,  
diphosphate, triphosphate, or a stabilized phosphate prodrug, H-  
phosphonate, including stabilized H-phosphonates, acyl, including  
optionally substituted phenyl and lower acyl, alkyl, including lower  
20 alkyl, O-substituted carboxyalkylamino or its peptide derivatives,

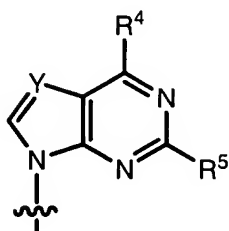


sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid, a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein  $R^1$  or  $R^7$  is independently H or phosphate;  $R^1$  and  $R^7$  can also be linked with cyclic phosphate group;

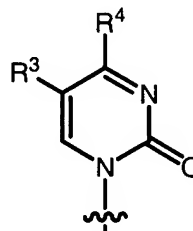
optionally, in a pharmaceutically acceptable carrier.

37. The method of claim 36, wherein

Base is selected from the group consisting of:



(a)



(b)

Y is N or CH;

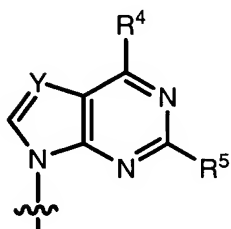
$R^3$ ,  $R^4$  and  $R^5$  are independently H, halogen including F, Cl, Br, I, OH, OR', SH, SR', NH<sub>2</sub>, NHR', NR'<sub>2</sub>, lower alkyl of C<sub>1</sub>-C<sub>6</sub>, halogenated (F, Cl, Br, I) lower alkyl of C<sub>1</sub>-C<sub>6</sub> such as CF<sub>3</sub> and CH<sub>2</sub>CH<sub>2</sub>F, lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CH<sub>2</sub>, halogenated (F, Cl, Br, I) lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CHCl, CH=CHBr and CH=CHI, lower alkynyl of C<sub>2</sub>-C<sub>6</sub> such as C≡CH, halogenated (F, Cl, Br, I) lower alkynyl of C<sub>2</sub>-C<sub>6</sub>, lower alkoxy of C<sub>1</sub>-C<sub>6</sub> such as CH<sub>2</sub>OH and CH<sub>2</sub>CH<sub>2</sub>OH, halogenated (F, Cl, Br, I) lower alkoxy of C<sub>1</sub>-C<sub>6</sub>, CO<sub>2</sub>H, CO<sub>2</sub>R', CONH<sub>2</sub>, CONHR', CONR'<sub>2</sub>, CH=CHCO<sub>2</sub>H, CH=CHCO<sub>2</sub>R'; and,

R' is an optionally substituted alkyl of C<sub>1</sub>-C<sub>12</sub> (particularly when the alkyl is an amino acid residue), cycloalkyl, optionally substituted alkynyl of C<sub>2</sub>-C<sub>6</sub>, optionally substituted lower alkenyl of C<sub>2</sub>-C<sub>6</sub>, or optionally substituted acyl.

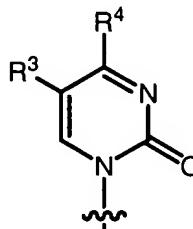
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38. The method of claim 36, wherein

Base is selected from the group consisting of (a) or (b):



(a)

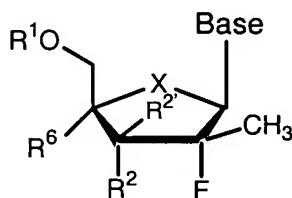


(b)

10 and wherein R<sup>1</sup> and R<sup>7</sup> are H, R<sup>3</sup> is H, and R<sup>4</sup> is NH<sub>2</sub> or OH, and R<sup>5</sup> is NH<sub>2</sub>.

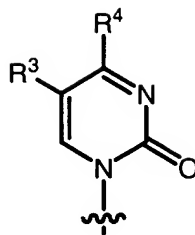
39. A method for the treatment or prophylaxis of hepatitis C infection comprising administering to a host an antivirally effective amount of a (2'R)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside (β-D or β-L) of the formula:

15



wherein

Base is



X is O, S, CH<sub>2</sub>, Se, NH, N-alkyl, CHW (R, S, or racemic), C(W)<sub>2</sub>, wherein W is F, Cl, Br, or I;

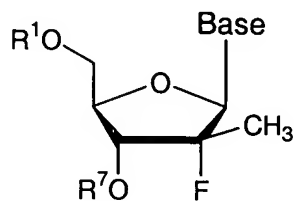
R<sup>1</sup> and R<sup>7</sup> are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid (or racemic mixture), a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein R<sup>1</sup> is H or phosphate; R<sup>2</sup> is H or phosphate; R<sup>1</sup> and R<sup>2</sup> or R<sup>7</sup> can also be linked with cyclic phosphate group;

R<sup>2</sup> and R<sup>2'</sup> are independently H, C<sub>1-4</sub> alkyl, C<sub>1-4</sub> alkenyl, C<sub>1-4</sub> alkynyl, vinyl, N<sub>3</sub>, CN, Cl, Br, F, I, NO<sub>2</sub>, C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkynyl), C(O)O(C<sub>1-4</sub> alkenyl), O(C<sub>1-4</sub> acyl), O(C<sub>1-4</sub> alkyl), O(C<sub>1-4</sub> alkenyl), S(C<sub>1-4</sub> acyl), S(C<sub>1-4</sub> alkyl), S(C<sub>1-4</sub> alkynyl), S(C<sub>1-4</sub> alkenyl), SO(C<sub>1-4</sub> acyl), SO(C<sub>1-4</sub> alkyl), SO(C<sub>1-4</sub> alkynyl), SO(C<sub>1-4</sub> alkenyl), SO<sub>2</sub>(C<sub>1-4</sub> acyl), SO<sub>2</sub>(C<sub>1-4</sub> alkyl), SO<sub>2</sub>(C<sub>1-4</sub> alkynyl), SO<sub>2</sub>(C<sub>1-4</sub> alkenyl), O<sub>3</sub>S(C<sub>1-4</sub> acyl), O<sub>3</sub>S(C<sub>1-4</sub> alkyl), O<sub>3</sub>S(C<sub>1-4</sub> alkenyl), NH<sub>2</sub>, NH(C<sub>1-4</sub> alkyl), NH(C<sub>1-4</sub> alkenyl), NH(C<sub>1-4</sub> alkynyl), NH(C<sub>1-4</sub> acyl), N(C<sub>1-4</sub> alkyl)<sub>2</sub>, N(C<sub>1-18</sub> acyl)<sub>2</sub>, wherein alkyl, alkynyl, alkenyl and vinyl are optionally substituted by N<sub>3</sub>, CN, one to three halogen (Cl, Br, F, I),

- 5 NO<sub>2</sub>, C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkynyl), C(O)O(C<sub>1-4</sub> alkenyl), O(C<sub>1-4</sub> acyl), O(C<sub>1-4</sub> alkyl), O(C<sub>1-4</sub> alkenyl), S(C<sub>1-4</sub> acyl), S(C<sub>1-4</sub> alkyl), S(C<sub>1-4</sub> alkynyl), S(C<sub>1-4</sub> alkenyl), SO(C<sub>1-4</sub> acyl), SO(C<sub>1-4</sub> alkyl), SO(C<sub>1-4</sub> alkynyl), SO(C<sub>1-4</sub> alkenyl), SO<sub>2</sub>(C<sub>1-4</sub> acyl), SO<sub>2</sub>(C<sub>1-4</sub> alkyl), SO<sub>2</sub>(C<sub>1-4</sub> alkynyl), SO<sub>2</sub>(C<sub>1-4</sub> alkenyl), O<sub>3</sub>S(C<sub>1-4</sub> acyl), O<sub>3</sub>S(C<sub>1-4</sub> alkyl), O<sub>3</sub>S(C<sub>1-4</sub> alkenyl), NH<sub>2</sub>, NH(C<sub>1-4</sub> alkyl), NH(C<sub>1-4</sub> alkenyl), NH(C<sub>1-4</sub> alkynyl), NH(C<sub>1-4</sub> acyl), N(C<sub>1-4</sub> alkyl)<sub>2</sub>, N(C<sub>1-4</sub> acyl)<sub>2</sub>, OR<sup>7</sup>; R<sup>2</sup> and R<sup>2'</sup> can be linked together to form a vinyl optionally substituted by one or two of N<sub>3</sub>, CN, Cl, Br, F, I, NO<sub>2</sub>;
- 10 R<sup>3</sup> and R<sup>4</sup> are independently H, halogen including F, Cl, Br, I, OH, OR', SH, SR', NH<sub>2</sub>, NHR', NR'<sub>2</sub>, lower alkyl of C<sub>1</sub>-C<sub>6</sub>, halogenated (F, Cl, Br, I) lower alkyl of C<sub>1</sub>-C<sub>6</sub> such as CF<sub>3</sub> and CH<sub>2</sub>CH<sub>2</sub>F, lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CH<sub>2</sub>, halogenated (F, Cl, Br, I) lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CHCl, CH=CHBr and CH=CHI, lower alkynyl of C<sub>2</sub>-C<sub>6</sub> such as C≡CH, halogenated (F, Cl, Br, I) lower alkynyl of C<sub>2</sub>-C<sub>6</sub>, lower alkoxy of C<sub>1</sub>-C<sub>6</sub> such as CH<sub>2</sub>OH and CH<sub>2</sub>CH<sub>2</sub>OH, halogenated (F, Cl, Br, I) lower alkoxy of C<sub>1</sub>-C<sub>6</sub>, CO<sub>2</sub>H, CO<sub>2</sub>R', CONH<sub>2</sub>, CONHR', CONR'<sub>2</sub>, CH=CHCO<sub>2</sub>H, CH=CHCO<sub>2</sub>R';
- 15
- 20 R' is an optionally substituted alkyl of C<sub>1</sub>-C<sub>12</sub> (particularly when the alkyl is an amino acid residue), cycloalkyl, optionally substituted alkynyl of C<sub>2</sub>-C<sub>6</sub>, optionally substituted lower alkenyl of C<sub>2</sub>-C<sub>6</sub>, or optionally substituted acyl; and,
- 25 R<sup>6</sup> is an optionally substituted alkyl (including lower alkyl), cyano (CN), CH<sub>3</sub>, OCH<sub>3</sub>, OCH<sub>2</sub>CH<sub>3</sub>, hydroxy methyl (CH<sub>2</sub>OH), fluoromethyl (CH<sub>2</sub>F), azido (N<sub>3</sub>), CHCN, CH<sub>2</sub>N<sub>3</sub>, CH<sub>2</sub>NH<sub>2</sub>, CH<sub>2</sub>NHCH<sub>3</sub>, CH<sub>2</sub>N(CH<sub>3</sub>)<sub>2</sub>, alkyne (optionally substituted), or fluoro;

or its pharmaceutically acceptable salt or prodrug thereof, optionally in a pharmaceutically acceptable carrier

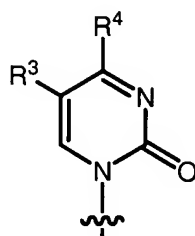
40. A method for the treatment or prophylaxis of hepatitis C infection comprising administering to a host an antivirally effective amount of a (2'R)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside ( $\beta$ -D or  $\beta$ -L) of the formula:



5

wherein

Base is



10

$R^1$  and  $R^7$  are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid, a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein  $R^1$  or  $R^7$  is independently H or phosphate;  $R^1$  and  $R^7$  can also be linked with cyclic phosphate group;

15

20

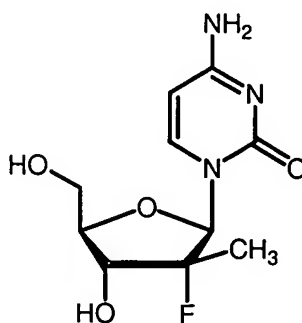
$R^3$  and  $R^4$  are independently H, halogen including F, Cl, Br, I, OH, OR', SH, SR', NH<sub>2</sub>, NHR', NR'<sub>2</sub>, lower alkyl of C<sub>1</sub>-C<sub>6</sub>, halogenated (F, Cl, Br, I)

lower alkyl of C<sub>1</sub>-C<sub>6</sub> such as CF<sub>3</sub> and CH<sub>2</sub>CH<sub>2</sub>F, lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CH<sub>2</sub>, halogenated (F, Cl, Br, I) lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CHCl, CH=CHBr and CH=CHI, lower alkynyl of C<sub>2</sub>-C<sub>6</sub> such as C≡CH, halogenated (F, Cl, Br, I) lower alkynyl of C<sub>2</sub>-C<sub>6</sub>, lower alkoxy of C<sub>1</sub>-C<sub>6</sub> such as CH<sub>2</sub>OH and CH<sub>2</sub>CH<sub>2</sub>OH, halogenated (F, Cl, Br, I) lower alkoxy of C<sub>1</sub>-C<sub>6</sub>, CO<sub>2</sub>H, CO<sub>2</sub>R', CONH<sub>2</sub>, CONHR', CONR'<sub>2</sub>, CH=CHCO<sub>2</sub>H, CH=CHCO<sub>2</sub>R'; and

R' is an optionally substituted alkyl of C<sub>1</sub>-C<sub>12</sub> (particularly when the alkyl is an amino acid residue), cycloalkyl, optionally substituted alkynyl of C<sub>2</sub>-C<sub>6</sub>, optionally substituted lower alkenyl of C<sub>2</sub>-C<sub>6</sub>, or optionally substituted acyl;

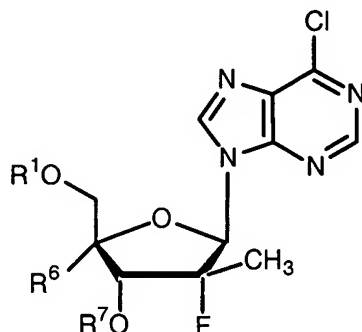
or its pharmaceutically acceptable salt or prodrug thereof, optionally in a pharmaceutically acceptable carrier

41. A method for the treatment or prophylaxis of hepatitis C infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside (β-D) or its pharmaceutically acceptable salt or prodrug thereof of the formula:



optionally in a pharmaceutically acceptable carrier.

42. A method for the treatment or prophylaxis of hepatitis C infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside ( $\beta$ -D) or its pharmaceutically acceptable salt or prodrug thereof of the formula:



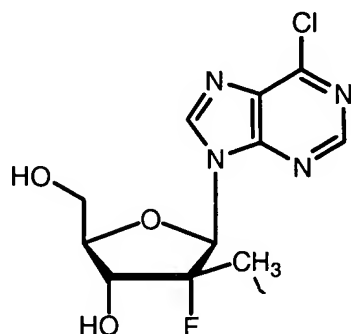
wherein

$R^1$  and  $R^7$  are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid, a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein  $R^1$  or  $R^7$  is independently H or phosphate;  $R^1$  and  $R^7$  can also be linked with cyclic phosphate group; and,

$R^6$  is an optionally substituted alkyl (including lower alkyl), cyano (CN),  $CH_3$ ,  $OCH_3$ ,  $OCH_2CH_3$ , hydroxy methyl ( $CH_2OH$ ), fluoromethyl ( $CH_2F$ ), azido ( $N_3$ ),  $CHCN$ ,  $CH_2N_3$ ,  $CH_2NH_2$ ,  $CH_2NHCH_3$ ,  $CH_2N(CH_3)_2$ , alkyne (optionally substituted), or fluoro;

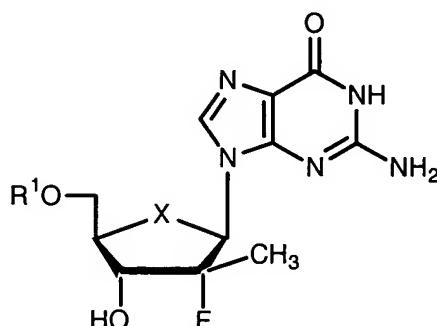
optionally in a pharmaceutically acceptable carrier.

43. A method for the treatment or prophylaxis of hepatitis C infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside ( $\beta$ -D) or its pharmaceutically acceptable salt or prodrug thereof of the formula:



optionally in a pharmaceutically acceptable carrier.

44. A method for the treatment or prophylaxis of hepatitis C infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside ( $\beta$ -D) or its pharmaceutically acceptable salt or prodrug thereof of the formula:



wherein

X is O, S, CH<sub>2</sub>, Se, NH, N-alkyl, CHW (*R*, *S*, or racemic), C(W)<sub>2</sub>, wherein W is F, Cl, Br, or I; and

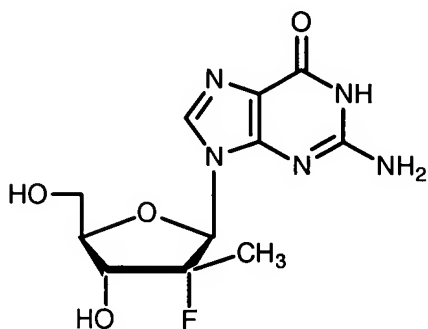
R<sup>1</sup> is H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-



phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid, a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein R<sup>1</sup> is H or phosphate;

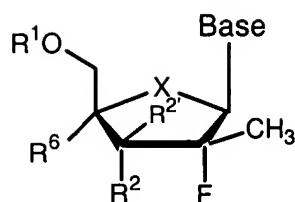
optionally in a pharmaceutically acceptable carrier.

45. A method for the treatment or prophylaxis of hepatitis C infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside ( $\beta$ -D) or its pharmaceutically acceptable salt or prodrug thereof of the formula:



optionally in a pharmaceutically acceptable carrier.

46. A method for the treatment or prophylaxis of a rhinovirus infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside ( $\beta$ -D or  $\beta$ -L) of the formula:



wherein

Base is a purine or pyrimidine base;

X is O, S, CH<sub>2</sub>, Se, NH, N-alkyl, CHW (*R*, *S*, or racemic), C(W)<sub>2</sub>, wherein W is F, Cl, Br, or I;

R<sup>1</sup> and R<sup>7</sup> are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid (or racemic mixture), a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein R<sup>1</sup> is H or phosphate; R<sup>2</sup> is H or phosphate; R<sup>1</sup> and R<sup>2</sup> or R<sup>7</sup> can also be linked with cyclic phosphate group;

R<sup>2</sup> and R<sup>2'</sup> are independently H, C<sub>1-4</sub> alkyl, C<sub>1-4</sub> alkenyl, C<sub>1-4</sub> alkynyl, vinyl, N<sub>3</sub>, CN, Cl, Br, F, I, NO<sub>2</sub>, C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkynyl), C(O)O(C<sub>1-4</sub> alkenyl), O(C<sub>1-4</sub> acyl), O(C<sub>1-4</sub> alkyl), O(C<sub>1-4</sub> alkenyl), S(C<sub>1-4</sub> acyl), S(C<sub>1-4</sub> alkyl), S(C<sub>1-4</sub> alkynyl), S(C<sub>1-4</sub> alkenyl), SO(C<sub>1-4</sub> acyl), SO(C<sub>1-4</sub> alkyl), SO(C<sub>1-4</sub> alkynyl), SO(C<sub>1-4</sub> alkenyl), SO<sub>2</sub>(C<sub>1-4</sub> acyl), SO<sub>2</sub>(C<sub>1-4</sub> alkyl), SO<sub>2</sub>(C<sub>1-4</sub> alkynyl), SO<sub>2</sub>(C<sub>1-4</sub> alkenyl), O<sub>3</sub>S(C<sub>1-4</sub> acyl), O<sub>3</sub>S(C<sub>1-4</sub> alkyl), O<sub>3</sub>S(C<sub>1-4</sub> alkenyl), NH<sub>2</sub>, NH(C<sub>1-4</sub> alkyl), NH(C<sub>1-4</sub> alkenyl), NH(C<sub>1-4</sub> alkynyl), NH(C<sub>1-4</sub> acyl), N(C<sub>1-4</sub>

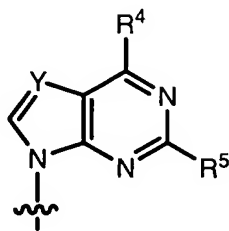
alkyl)<sub>2</sub>, N(C<sub>1-18</sub> acyl)<sub>2</sub>, wherein alkyl, alkynyl, alkenyl and vinyl are optionally substituted by N<sub>3</sub>, CN, one to three halogen (Cl, Br, F, I), NO<sub>2</sub>, C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkynyl), C(O)O(C<sub>1-4</sub> alkenyl), O(C<sub>1-4</sub> acyl), O(C<sub>1-4</sub> alkyl), O(C<sub>1-4</sub> alkenyl), S(C<sub>1-4</sub> acyl), S(C<sub>1-4</sub> alkyl), S(C<sub>1-4</sub> alkynyl), S(C<sub>1-4</sub> alkenyl), SO(C<sub>1-4</sub> acyl), SO(C<sub>1-4</sub> alkyl), SO(C<sub>1-4</sub> alkynyl), SO(C<sub>1-4</sub> alkenyl), SO<sub>2</sub>(C<sub>1-4</sub> acyl), SO<sub>2</sub>(C<sub>1-4</sub> alkyl), SO<sub>2</sub>(C<sub>1-4</sub> alkynyl), SO<sub>2</sub>(C<sub>1-4</sub> alkenyl), O<sub>3</sub>S(C<sub>1-4</sub> acyl), O<sub>3</sub>S(C<sub>1-4</sub> alkyl), O<sub>3</sub>S(C<sub>1-4</sub> alkenyl), NH<sub>2</sub>, NH(C<sub>1-4</sub> alkyl), NH(C<sub>1-4</sub> alkenyl), NH(C<sub>1-4</sub> alkynyl), NH(C<sub>1-4</sub> acyl), N(C<sub>1-4</sub> alkyl)<sub>2</sub>, N(C<sub>1-4</sub> acyl)<sub>2</sub>, OR<sup>7</sup>; R<sup>2</sup> and R<sup>2'</sup> can be linked together to form a vinyl optionally substituted by one or two of N<sub>3</sub>, CN, Cl, Br, F, I, NO<sub>2</sub>;

R<sup>6</sup> is an optionally substituted alkyl (including lower alkyl), cyano (CN), CH<sub>3</sub>, OCH<sub>3</sub>, OCH<sub>2</sub>CH<sub>3</sub>, hydroxy methyl (CH<sub>2</sub>OH), fluoromethyl (CH<sub>2</sub>F), azido (N<sub>3</sub>), CHCN, CH<sub>2</sub>N<sub>3</sub>, CH<sub>2</sub>NH<sub>2</sub>, CH<sub>2</sub>NHCH<sub>3</sub>, CH<sub>2</sub>N(CH<sub>3</sub>)<sub>2</sub>, alkyne (optionally substituted), or fluoro;

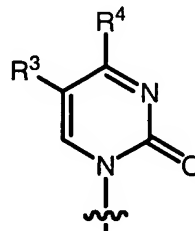
or its pharmaceutically acceptable salt or prodrug thereof, optionally in a pharmaceutically acceptable carrier.

47. The method of claim 46,

wherein Base is selected from the group consisting of:



(a)



(b)

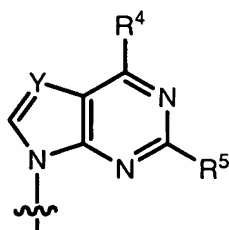
Y is N or CH.

$R^3$ ,  $R^4$  and  $R^5$  are independently H, halogen including F, Cl, Br, I, OH, OR', SH, SR', NH<sub>2</sub>, NHR', NR'<sub>2</sub>, lower alkyl of C<sub>1</sub>-C<sub>6</sub>, halogenated (F, Cl, Br, I) lower alkyl of C<sub>1</sub>-C<sub>6</sub> such as CF<sub>3</sub> and CH<sub>2</sub>CH<sub>2</sub>F, lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CH<sub>2</sub>, halogenated (F, Cl, Br, I) lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CHCl, CH=CHBr and CH=CHI, lower alkynyl of C<sub>2</sub>-C<sub>6</sub> such as C≡CH, halogenated (F, Cl, Br, I) lower alkynyl of C<sub>2</sub>-C<sub>6</sub>, lower alkoxy of C<sub>1</sub>-C<sub>6</sub> such as CH<sub>2</sub>OH and CH<sub>2</sub>CH<sub>2</sub>OH, halogenated (F, Cl, Br, I) lower alkoxy of C<sub>1</sub>-C<sub>6</sub>, CO<sub>2</sub>H, CO<sub>2</sub>R', CONH<sub>2</sub>, CONHR', CONR'<sub>2</sub>, CH=CHCO<sub>2</sub>H, CH=CHCO<sub>2</sub>R'; and,

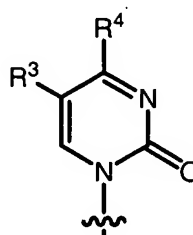
R' is an optionally substituted alkyl of C<sub>1</sub>-C<sub>12</sub> (particularly when the alkyl is an amino acid residue), cycloalkyl, optionally substituted alkynyl of C<sub>2</sub>-C<sub>6</sub>, optionally substituted lower alkenyl of C<sub>2</sub>-C<sub>6</sub>, or optionally substituted acyl.

48. The method of claim 46, wherein

Base is selected from the group consisting of (a) or (b):



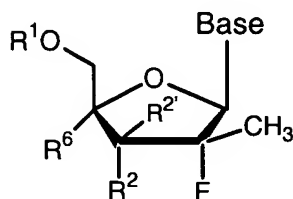
(a)



(b)

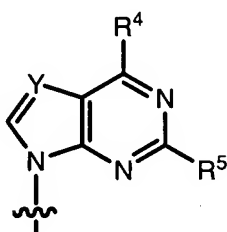
and wherein  $R^1$  is H,  $R^2$  is OH,  $R^{2'}$  is H,  $R^3$  is H, and  $R^4$  is NH<sub>2</sub> or OH, and  $R^5$  is NH<sub>2</sub>.

49. A method for the treatment or prophylaxis of a rhinovirus infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside ( $\beta$ -D or  $\beta$ -L) of the formula:

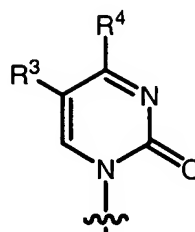


wherein

Base is selected from the group consisting of



(a)



(b)

Y is N or CH;

$R^1$  and  $R^7$  are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid (or racemic mixture), a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein  $R^1$  is H or phosphate;

$R^2$  is H or phosphate;  $R^1$  and  $R^2$  or  $R^7$  can also be linked with cyclic phosphate group;

$R^2$  and  $R^{2'}$  are independently H,  $C_{1-4}$  alkyl,  $C_{1-4}$  alkenyl,  $C_{1-4}$  alkynyl, vinyl,  $N_3$ , CN, Cl, Br, F, I,  $NO_2$ ,  $C(O)O(C_{1-4}$  alkyl),  $C(O)O(C_{1-4}$  alkenyl),  $C(O)O(C_{1-4}$  alkynyl),  $C(O)O(C_{1-4}$  acyl),  $O(C_{1-4}$  alkyl),  $O(C_{1-4}$  alkenyl),  $S(C_{1-4}$  alkyl),  $S(C_{1-4}$  alkenyl),  $SO(C_{1-4}$  alkyl),  $SO(C_{1-4}$  alkenyl),  $SO_2(C_{1-4}$  alkyl),  $SO_2(C_{1-4}$  alkenyl),  $O_3S(C_{1-4}$  alkyl),  $O_3S(C_{1-4}$  alkenyl),  $NH_2$ ,  $NH(C_{1-4}$  alkyl),  $NH(C_{1-4}$  alkenyl),  $NH(C_{1-4}$  alkynyl),  $NH(C_{1-4}$  acyl),  $N(C_{1-4}$  alkyl)<sub>2</sub>,  $N(C_{1-18}$  acyl)<sub>2</sub>, wherein alkyl, alkynyl, alkenyl and vinyl are optionally substituted by  $N_3$ , CN, one to three halogen (Cl, Br, F, I),  $NO_2$ ,  $C(O)O(C_{1-4}$  alkyl),  $C(O)O(C_{1-4}$  alkenyl),  $C(O)O(C_{1-4}$  alkynyl),  $C(O)O(C_{1-4}$  acyl),  $O(C_{1-4}$  alkyl),  $O(C_{1-4}$  alkenyl),  $S(C_{1-4}$  alkyl),  $S(C_{1-4}$  alkenyl),  $SO(C_{1-4}$  alkyl),  $SO(C_{1-4}$  alkenyl),  $SO_2(C_{1-4}$  alkyl),  $SO_2(C_{1-4}$  alkenyl),  $O_3S(C_{1-4}$  alkyl),  $O_3S(C_{1-4}$  alkenyl),  $NH_2$ ,  $NH(C_{1-4}$  alkyl),  $NH(C_{1-4}$  alkenyl),  $NH(C_{1-4}$  alkynyl),  $NH(C_{1-4}$  acyl),  $N(C_{1-4}$  alkyl)<sub>2</sub>,  $N(C_{1-4}$  acyl)<sub>2</sub>,  $OR^7$ ;  $R^2$  and  $R^{2'}$  can be linked together to form a vinyl optionally substituted by one or two of  $N_3$ , CN, Cl, Br, F, I,  $NO_2$ ;

$R^3$ ,  $R^4$  and  $R^5$  are independently H, halogen including F, Cl, Br, I, OH,  $OR'$ , SH,  $SR'$ ,  $NH_2$ ,  $NHR'$ ,  $NR'_2$ , lower alkyl of  $C_1-C_6$ , halogenated (F, Cl, Br, I) lower alkyl of  $C_1-C_6$  such as  $CF_3$  and  $CH_2CH_2F$ , lower alkenyl of  $C_2-C_6$  such as  $CH=CH_2$ , halogenated (F, Cl, Br, I) lower alkenyl of  $C_2-C_6$  such as  $CH=CHCl$ ,  $CH=CHBr$  and  $CH=CHI$ , lower alkynyl of  $C_2-C_6$  such as  $C\equiv CH$ , halogenated (F, Cl, Br, I) lower alkynyl of  $C_2-C_6$ , lower alkoxy of  $C_1-C_6$  such as  $CH_2OH$  and  $CH_2CH_2OH$ , halogenated (F, Cl, Br, I) lower alkoxy of  $C_1-C_6$ ,  $CO_2H$ ,  $CO_2R'$ ,  $CONH_2$ ,  $CONHR'$ ,  $CONR'_2$ ,  $CH=CHCO_2H$ ,  $CH=CHCO_2R'$ ;

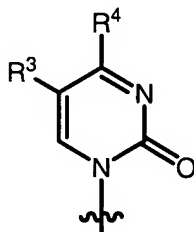
R' is an optionally substituted alkyl of C<sub>1</sub>-C<sub>12</sub> (particularly when the alkyl is an amino acid residue), cycloalkyl, optionally substituted alkynyl of C<sub>2</sub>-C<sub>6</sub>, optionally substituted lower alkenyl of C<sub>2</sub>-C<sub>6</sub>, or optionally substituted acyl;

5 R<sup>6</sup> is an optionally substituted alkyl (including lower alkyl), cyano (CN), CH<sub>3</sub>, OCH<sub>3</sub>, OCH<sub>2</sub>CH<sub>3</sub>, hydroxy methyl (CH<sub>2</sub>OH), fluoromethyl (CH<sub>2</sub>F), azido (N<sub>3</sub>), CHCN, CH<sub>2</sub>N<sub>3</sub>, CH<sub>2</sub>NH<sub>2</sub>, CH<sub>2</sub>NHCH<sub>3</sub>, CH<sub>2</sub>N(CH<sub>3</sub>)<sub>2</sub>, alkyne (optionally substituted), or fluoro;

10 or its pharmaceutically acceptable salt or prodrug thereof, optionally in a pharmaceutically acceptable carrier.

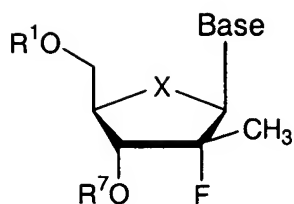
50. The method of claim 49, wherein

Base is



15 and R<sup>1</sup> is H, R<sup>2</sup> is OH, R<sup>2'</sup> is H, R<sup>3</sup> is H, R<sup>4</sup> is NH<sub>2</sub> or OH, and R<sup>6</sup> is H.

20 51. A method for the treatment or prophylaxis of a rhinovirus infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside (β-D or β-L) or its pharmaceutically acceptable salt or prodrug thereof of the structure:



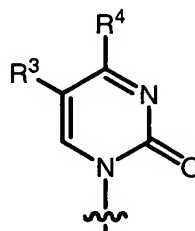
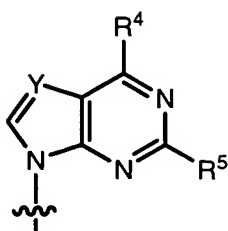
wherein Base is a purine or pyrimidine base;

X is O, S, CH<sub>2</sub>, Se, NH, N-alkyl, CHW (*R*, *S*, or racemic), C(W)<sub>2</sub>, wherein W is F, Cl, Br, or I; and,

5           R<sup>1</sup> and R<sup>7</sup> are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, 10           sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid, a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of 15           providing a compound wherein R<sup>1</sup> or R<sup>7</sup> is independently H or phosphate; R<sup>1</sup> and R<sup>7</sup> can also be linked with cyclic phosphate group and optionally a pharmaceutically acceptable carrier.

52.   The method of claim 51, wherein

20           Base is selected from the group consisting of:





(a)

(b)

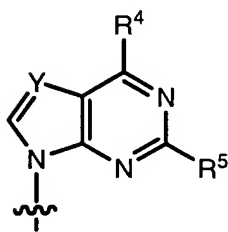
Y is N or CH;

$R^3$ ,  $R^4$  and  $R^5$  are independently H, halogen including F, Cl, Br, I, OH, OR', SH, SR', NH<sub>2</sub>, NHR', NR'<sub>2</sub>, lower alkyl of C<sub>1</sub>-C<sub>6</sub>, halogenated (F, Cl, Br, I) lower alkyl of C<sub>1</sub>-C<sub>6</sub> such as CF<sub>3</sub> and CH<sub>2</sub>CH<sub>2</sub>F, lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CH<sub>2</sub>, halogenated (F, Cl, Br, I) lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CHCl, CH=CHBr and CH=CHI, lower alkynyl of C<sub>2</sub>-C<sub>6</sub> such as C≡CH, halogenated (F, Cl, Br, I) lower alkynyl of C<sub>2</sub>-C<sub>6</sub>, lower alkoxy of C<sub>1</sub>-C<sub>6</sub> such as CH<sub>2</sub>OH and CH<sub>2</sub>CH<sub>2</sub>OH, halogenated (F, Cl, Br, I) lower alkoxy of C<sub>1</sub>-C<sub>6</sub>, CO<sub>2</sub>H, CO<sub>2</sub>R', CONH<sub>2</sub>, CONHR', CONR'<sub>2</sub>, CH=CHCO<sub>2</sub>H, CH=CHCO<sub>2</sub>R'; and,

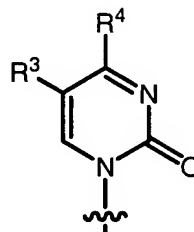
R' is an optionally substituted alkyl of C<sub>1</sub>-C<sub>12</sub> (particularly when the alkyl is an amino acid residue), cycloalkyl, optionally substituted alkynyl of C<sub>2</sub>-C<sub>6</sub>, optionally substituted lower alkenyl of C<sub>2</sub>-C<sub>6</sub>, or optionally substituted acyl.

53. The method of claim 51, wherein

Base is selected from the group consisting of (a) or (b):



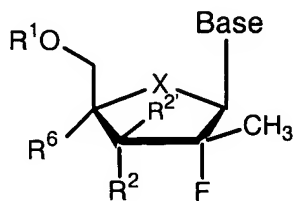
(a)



(b)

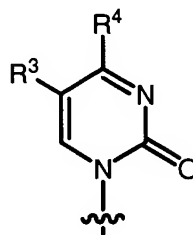
and wherein R<sup>1</sup> and R<sup>7</sup> are H, R<sup>3</sup> is H, and R<sup>4</sup> is NH<sub>2</sub> or OH, and R<sup>5</sup> is NH<sub>2</sub>.

54. A method for the treatment or prophylaxis of a rhinovirus infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside ( $\beta$ -D or  $\beta$ -L) of the formula:



wherein

Base is



X is O, S, CH<sub>2</sub>, Se, NH, N-alkyl, CHW (*R*, *S*, or racemic), C(W)<sub>2</sub>, wherein W is F, Cl, Br, or I;

$R^1$  and  $R^7$  are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid (or racemic mixture), a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein  $R^1$  is H or phosphate;  $R^2$  is H or phosphate;  $R^1$  and  $R^2$  or  $R^7$  can also be linked with cyclic phosphate group;

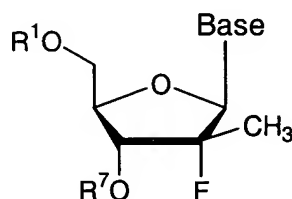
$R^2$  and  $R^{2'}$  are independently H,  $C_{1-4}$  alkyl,  $C_{1-4}$  alkenyl,  $C_{1-4}$  alkynyl, vinyl,  $N_3$ ,  
 CN, Cl, Br, F, I,  $NO_2$ ,  $C(O)O(C_{1-4} \text{ alkyl})$ ,  $C(O)O(C_{1-4} \text{ alkenyl})$ ,  $C(O)O(C_{1-4} \text{ alkynyl})$ ,  
 5  $C(O)O(C_{1-4} \text{ alkenyl})$ ,  $O(C_{1-4} \text{ acyl})$ ,  $O(C_{1-4} \text{ alkyl})$ ,  $O(C_{1-4} \text{ alkenyl})$ ,  $S(C_{1-4} \text{ acyl})$ ,  
 $S(C_{1-4} \text{ alkyl})$ ,  $S(C_{1-4} \text{ alkynyl})$ ,  $S(C_{1-4} \text{ alkenyl})$ ,  $SO(C_{1-4} \text{ acyl})$ ,  $SO(C_{1-4} \text{ alkyl})$ ,  
 $SO(C_{1-4} \text{ alkynyl})$ ,  $SO(C_{1-4} \text{ alkenyl})$ ,  $SO_2(C_{1-4} \text{ acyl})$ ,  $SO_2(C_{1-4} \text{ alkyl})$ ,  
 $SO_2(C_{1-4} \text{ alkynyl})$ ,  $SO_2(C_{1-4} \text{ alkenyl})$ ,  $O_3S(C_{1-4} \text{ acyl})$ ,  $O_3S(C_{1-4} \text{ alkyl})$ ,  
 $O_3S(C_{1-4} \text{ alkenyl})$ ,  $NH_2$ ,  $NH(C_{1-4} \text{ alkyl})$ ,  $NH(C_{1-4} \text{ alkenyl})$ ,  $NH(C_{1-4} \text{ alkynyl})$ ,  
 10  $NH(C_{1-4} \text{ acyl})$ ,  $N(C_{1-4} \text{ alkyl})_2$ ,  $N(C_{1-18} \text{ acyl})_2$ , wherein alkyl, alkynyl, alkenyl and vinyl are  
 optionally substituted by  $N_3$ , CN, one to three halogen (Cl, Br, F, I),  $NO_2$ ,  $C(O)O(C_{1-4} \text{ alkyl})$ ,  
 $C(O)O(C_{1-4} \text{ alkenyl})$ ,  $C(O)O(C_{1-4} \text{ alkynyl})$ ,  $C(O)O(C_{1-4} \text{ alkenyl})$ ,  $O(C_{1-4} \text{ acyl})$ ,  
 $O(C_{1-4} \text{ alkyl})$ ,  $O(C_{1-4} \text{ alkenyl})$ ,  $S(C_{1-4} \text{ acyl})$ ,  $S(C_{1-4} \text{ alkyl})$ ,  $S(C_{1-4} \text{ alkynyl})$ ,  
 15  $S(C_{1-4} \text{ alkenyl})$ ,  $SO(C_{1-4} \text{ acyl})$ ,  $SO(C_{1-4} \text{ alkyl})$ ,  $SO(C_{1-4} \text{ alkynyl})$ ,  $SO(C_{1-4} \text{ alkenyl})$ ,  
 $SO_2(C_{1-4} \text{ acyl})$ ,  $SO_2(C_{1-4} \text{ alkyl})$ ,  $SO_2(C_{1-4} \text{ alkynyl})$ ,  $SO_2(C_{1-4} \text{ alkenyl})$ ,  
 $O_3S(C_{1-4} \text{ acyl})$ ,  $O_3S(C_{1-4} \text{ alkyl})$ ,  $O_3S(C_{1-4} \text{ alkenyl})$ ,  $NH_2$ ,  $NH(C_{1-4} \text{ alkyl})$ ,  
 $NH(C_{1-4} \text{ alkenyl})$ ,  $NH(C_{1-4} \text{ alkynyl})$ ,  $NH(C_{1-4} \text{ acyl})$ ,  $N(C_{1-4} \text{ alkyl})_2$ ,  
 $N(C_{1-4} \text{ acyl})_2$ ,  $OR^7$ ;  $R^2$  and  $R^{2'}$  can be linked together to form a vinyl optionally  
 substituted by one or two of  $N_3$ , CN, Cl, Br, F, I,  $NO_2$ ;  
 20  $R^3$  and  $R^4$  are independently H, halogen including F, Cl, Br, I, OH,  $OR'$ , SH,  
 $SR'$ ,  $NH_2$ ,  $NHR'$ ,  $NR'_2$ , lower alkyl of  $C_1-C_6$ , halogenated (F, Cl, Br, I)  
 lower alkyl of  $C_1-C_6$  such as  $CF_3$  and  $CH_2CH_2F$ , lower alkenyl of  $C_2-C_6$   
 such as  $CH=CH_2$ , halogenated (F, Cl, Br, I) lower alkenyl of  $C_2-C_6$   
 such as  $CH=CHCl$ ,  $CH=CHBr$  and  $CH=CHI$ , lower alkynyl of  $C_2-C_6$   
 25 such as  $C\equiv CH$ , halogenated (F, Cl, Br, I) lower alkynyl of  $C_2-C_6$ , lower  
 alkoxy of  $C_1-C_6$  such as  $CH_2OH$  and  $CH_2CH_2OH$ , halogenated (F, Cl,  
 Br, I) lower alkoxy of  $C_1-C_6$ ,  $CO_2H$ ,  $CO_2R'$ ,  $CONH_2$ ,  $CONHR'$ ,  
 $CONR'_2$ ,  $CH=CHCO_2H$ ,  $CH=CHCO_2R'$ ;  
 30  $R'$  is an optionally substituted alkyl of  $C_1-C_{12}$  (particularly when the alkyl is an  
 amino acid residue), cycloalkyl, optionally substituted alkynyl of  $C_2-$

C<sub>6</sub>, optionally substituted lower alkenyl of C<sub>2</sub>-C<sub>6</sub>, or optionally substituted acyl;

R<sup>6</sup> is an optionally substituted alkyl (including lower alkyl), cyano (CN), CH<sub>3</sub>, OCH<sub>3</sub>, OCH<sub>2</sub>CH<sub>3</sub>, hydroxy methyl (CH<sub>2</sub>OH), fluoromethyl (CH<sub>2</sub>F), azido (N<sub>3</sub>), CHCN, CH<sub>2</sub>N<sub>3</sub>, CH<sub>2</sub>NH<sub>2</sub>, CH<sub>2</sub>NHCH<sub>3</sub>, CH<sub>2</sub>N(CH<sub>3</sub>)<sub>2</sub>, alkyne (optionally substituted), or fluoro;

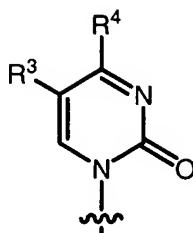
or its pharmaceutically acceptable salt or prodrug thereof, optionally in a pharmaceutically acceptable carrier.

55. A method for the treatment or prophylaxis of a rhinovirus infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside (β-D or β-L) of the formula:



wherein

Base is



R<sup>1</sup> and R<sup>7</sup> are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including

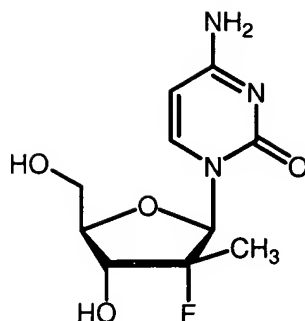
5 optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid, a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein  $R^1$  or  $R^7$  is independently H or phosphate;  $R^1$  and  $R^7$  can also be linked with cyclic phosphate group;

10  $R^3$  and  $R^4$  are independently H, halogen including F, Cl, Br, I, OH, OR', SH, SR', NH<sub>2</sub>, NHR', NR'<sub>2</sub>, lower alkyl of C<sub>1</sub>-C<sub>6</sub>, halogenated (F, Cl, Br, I) lower alkyl of C<sub>1</sub>-C<sub>6</sub> such as CF<sub>3</sub> and CH<sub>2</sub>CH<sub>2</sub>F, lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CH<sub>2</sub>, halogenated (F, Cl, Br, I) lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CHCl, CH=CHBr and CH=CHI, lower alkynyl of C<sub>2</sub>-C<sub>6</sub> such as C≡CH, halogenated (F, Cl, Br, I) lower alkynyl of C<sub>2</sub>-C<sub>6</sub>, lower alkoxy of C<sub>1</sub>-C<sub>6</sub> such as CH<sub>2</sub>OH and CH<sub>2</sub>CH<sub>2</sub>OH, halogenated (F, Cl, Br, I) lower alkoxy of C<sub>1</sub>-C<sub>6</sub>, CO<sub>2</sub>H, CO<sub>2</sub>R', CONH<sub>2</sub>, CONHR', CONR'<sub>2</sub>, CH=CHCO<sub>2</sub>H, CH=CHCO<sub>2</sub>R';

15  
20  $R'$  is an optionally substituted alkyl of C<sub>1</sub>-C<sub>12</sub> (particularly when the alkyl is an amino acid residue), cycloalkyl, optionally substituted alkynyl of C<sub>2</sub>-C<sub>6</sub>, optionally substituted lower alkenyl of C<sub>2</sub>-C<sub>6</sub>, or optionally substituted acyl;

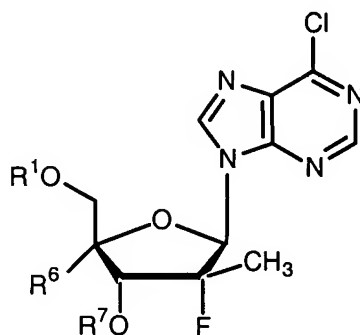
or its pharmaceutically acceptable salt or prodrug thereof, optionally in a pharmaceutically acceptable carrier.

25  
56. A method for the treatment or prophylaxis of a rhinovirus infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside (β-D) or its pharmaceutically acceptable salt or prodrug thereof of the formula:



optionally in a pharmaceutically acceptable carrier.

57. A method for the treatment or prophylaxis of a rhinovirus infection comprising administering to a host an antivirally effective amount of a (2*R*)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside (β-D) or its pharmaceutically acceptable salt or prodrug thereof of the formula:



wherein

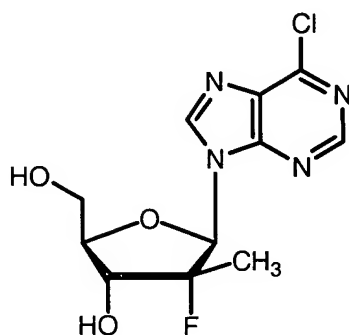
R<sup>1</sup> and R<sup>7</sup> are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid, a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of

providing a compound wherein  $R^1$  or  $R^7$  is independently H or phosphate;  $R^1$  and  $R^7$  can also be linked with cyclic phosphate group; and,

$R^6$  is an optionally substituted alkyl (including lower alkyl), cyano (CN),  $CH_3$ ,  $OCH_3$ ,  $OCH_2CH_3$ , hydroxy methyl ( $CH_2OH$ ), fluoromethyl ( $CH_2F$ ), azido ( $N_3$ ),  $CHCN$ ,  $CH_2N_3$ ,  $CH_2NH_2$ ,  $CH_2NHCH_3$ ,  $CH_2N(CH_3)_2$ , alkyne (optionally substituted), or fluoro; and

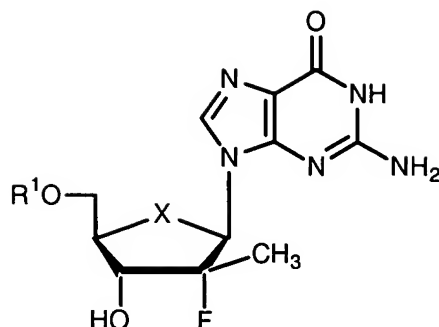
optionally in a pharmaceutically acceptable carrier.

58. A method for the treatment or prophylaxis of a rhinovirus infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside ( $\beta$ -D) or its pharmaceutically acceptable salt or prodrug thereof of the formula:



optionally in a pharmaceutically acceptable carrier.

59. A method for the treatment or prophylaxis of a rhinovirus infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside ( $\beta$ -D) or its pharmaceutically acceptable salt or prodrug thereof of the formula:



wherein

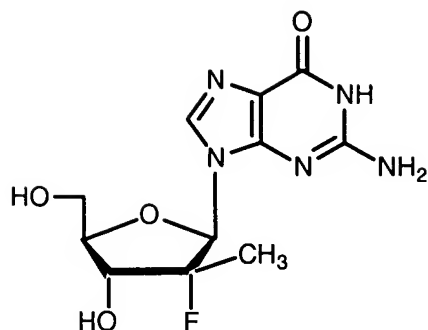
X is O, S, CH<sub>2</sub>, Se, NH, N-alkyl, CHW (*R*, *S*, or racemic), C(W)<sub>2</sub>, wherein W is F, Cl, Br, or I; and

- 5           R<sup>1</sup> is H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl
- 10           sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid, a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein R<sup>1</sup> is H or phosphate;
- 15           optionally in a pharmaceutically acceptable carrier.

60.     A method for the treatment or prophylaxis of a rhinovirus infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside (β-D) or its pharmaceutically acceptable salt or prodrug thereof of the

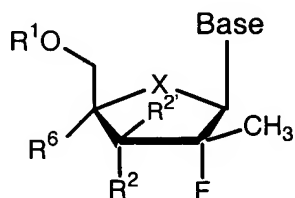
20           formula:





optionally in a pharmaceutically acceptable carrier.

61. A method for the treatment or prophylaxis of a yellow fever virus infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside ( $\beta$ -D or  $\beta$ -L) of the formula:



wherein

Base is a purine or pyrimidine base;

X is O, S, CH<sub>2</sub>, Se, NH, N-alkyl, CHW (*R*, *S*, or racemic), C(W)<sub>2</sub>, wherein W is F, Cl, Br, or I;

R<sup>1</sup> and R<sup>7</sup> are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid (or racemic mixture), a carbohydrate, a peptide, a cholesterol, or other

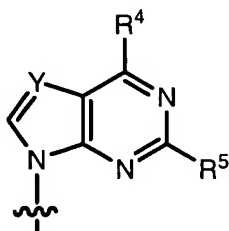
pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein R<sup>1</sup> is H or phosphate; R<sup>2</sup> is H or phosphate; R<sup>1</sup> and R<sup>2</sup> or R<sup>7</sup> can also be linked with cyclic phosphate group;

- 5 R<sup>2</sup> and R<sup>2'</sup> are independently H, C<sub>1-4</sub> alkyl, C<sub>1-4</sub> alkenyl, C<sub>1-4</sub> alkynyl, vinyl, N<sub>3</sub>, CN, Cl, Br, F, I, NO<sub>2</sub>, C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkynyl), C(O)O(C<sub>1-4</sub> alkenyl), O(C<sub>1-4</sub> acyl), O(C<sub>1-4</sub> alkyl), O(C<sub>1-4</sub> alkenyl), S(C<sub>1-4</sub> acyl), S(C<sub>1-4</sub> alkyl), S(C<sub>1-4</sub> alkynyl), S(C<sub>1-4</sub> alkenyl), SO(C<sub>1-4</sub> acyl), SO(C<sub>1-4</sub> alkyl), SO(C<sub>1-4</sub> alkynyl), SO(C<sub>1-4</sub> alkenyl), SO<sub>2</sub>(C<sub>1-4</sub> acyl), SO<sub>2</sub>(C<sub>1-4</sub> alkyl), SO<sub>2</sub>(C<sub>1-4</sub> alkynyl), SO<sub>2</sub>(C<sub>1-4</sub> alkenyl), O<sub>3</sub>S(C<sub>1-4</sub> acyl), O<sub>3</sub>S(C<sub>1-4</sub> alkyl), O<sub>3</sub>S(C<sub>1-4</sub> alkenyl), NH<sub>2</sub>, NH(C<sub>1-4</sub> alkyl), NH(C<sub>1-4</sub> alkenyl), NH(C<sub>1-4</sub> alkynyl), NH(C<sub>1-4</sub> acyl), N(C<sub>1-4</sub> alkyl)<sub>2</sub>, N(C<sub>1-18</sub> acyl)<sub>2</sub>, wherein alkyl, alkynyl, alkenyl and vinyl are optionally substituted by N<sub>3</sub>, CN, one to three halogen (Cl, Br, F, I), NO<sub>2</sub>, C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkynyl), C(O)O(C<sub>1-4</sub> alkenyl), O(C<sub>1-4</sub> acyl), O(C<sub>1-4</sub> alkyl), O(C<sub>1-4</sub> alkenyl), S(C<sub>1-4</sub> acyl), S(C<sub>1-4</sub> alkyl), S(C<sub>1-4</sub> alkynyl), S(C<sub>1-4</sub> alkenyl), SO(C<sub>1-4</sub> acyl), SO(C<sub>1-4</sub> alkyl), SO(C<sub>1-4</sub> alkynyl), SO(C<sub>1-4</sub> alkenyl), SO<sub>2</sub>(C<sub>1-4</sub> acyl), SO<sub>2</sub>(C<sub>1-4</sub> alkyl), SO<sub>2</sub>(C<sub>1-4</sub> alkynyl), SO<sub>2</sub>(C<sub>1-4</sub> alkenyl), O<sub>3</sub>S(C<sub>1-4</sub> acyl), O<sub>3</sub>S(C<sub>1-4</sub> alkyl), O<sub>3</sub>S(C<sub>1-4</sub> alkenyl), NH<sub>2</sub>, NH(C<sub>1-4</sub> alkyl), NH(C<sub>1-4</sub> alkenyl), NH(C<sub>1-4</sub> alkynyl), NH(C<sub>1-4</sub> acyl), N(C<sub>1-4</sub> alkyl)<sub>2</sub>, N(C<sub>1-4</sub> acyl)<sub>2</sub>, OR<sup>7</sup>; R<sup>2</sup> and R<sup>2'</sup> can be linked together to form a vinyl optionally substituted by one or two of N<sub>3</sub>, CN, Cl, Br, F, I, NO<sub>2</sub>;
- 10
- 15
- 20
- 25 R<sup>6</sup> is an optionally substituted alkyl (including lower alkyl), cyano (CN), CH<sub>3</sub>, OCH<sub>3</sub>, OCH<sub>2</sub>CH<sub>3</sub>, hydroxy methyl (CH<sub>2</sub>OH), fluoromethyl (CH<sub>2</sub>F), azido (N<sub>3</sub>), CHCN, CH<sub>2</sub>N<sub>3</sub>, CH<sub>2</sub>NH<sub>2</sub>, CH<sub>2</sub>NHCH<sub>3</sub>, CH<sub>2</sub>N(CH<sub>3</sub>)<sub>2</sub>, alkyne (optionally substituted), or fluoro;

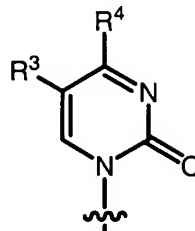
or its pharmaceutically acceptable salt or prodrug thereof, optionally in a pharmaceutically acceptable carrier.

62. The method of claim 61,

wherein Base is selected from the group consisting of:



(a)



(b)

5

Y is N or CH.

R<sup>3</sup>, R<sup>4</sup> and R<sup>5</sup> are independently H, halogen including F, Cl, Br, I, OH, OR', SH, SR', NH<sub>2</sub>, NHR', NR'<sub>2</sub>, lower alkyl of C<sub>1</sub>-C<sub>6</sub>, halogenated (F, Cl, Br, I) lower alkyl of C<sub>1</sub>-C<sub>6</sub> such as CF<sub>3</sub> and CH<sub>2</sub>CH<sub>2</sub>F, lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CH<sub>2</sub>, halogenated (F, Cl, Br, I) lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CHCl, CH=CHBr and CH=CHI, lower alkynyl of C<sub>2</sub>-C<sub>6</sub> such as C≡CH, halogenated (F, Cl, Br, I) lower alkynyl of C<sub>2</sub>-C<sub>6</sub>, lower alkoxy of C<sub>1</sub>-C<sub>6</sub> such as CH<sub>2</sub>OH and CH<sub>2</sub>CH<sub>2</sub>OH, halogenated (F, Cl, Br, I) lower alkoxy of C<sub>1</sub>-C<sub>6</sub>, CO<sub>2</sub>H, CO<sub>2</sub>R', CONH<sub>2</sub>, CONHR', CONR'<sub>2</sub>, CH=CHCO<sub>2</sub>H, CH=CHCO<sub>2</sub>R'; and,

10

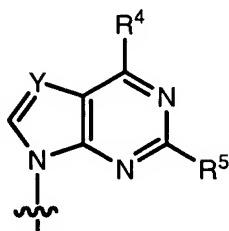
15

R' is an optionally substituted alkyl of C<sub>1</sub>-C<sub>12</sub> (particularly when the alkyl is an amino acid residue), cycloalkyl, optionally substituted alkynyl of C<sub>2</sub>-C<sub>6</sub>, optionally substituted lower alkenyl of C<sub>2</sub>-C<sub>6</sub>, or optionally substituted acyl.

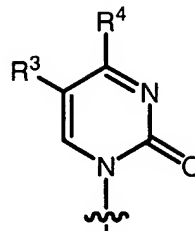
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63. The method of claim 61, wherein

Base is selected from the group consisting of (a) or (b):



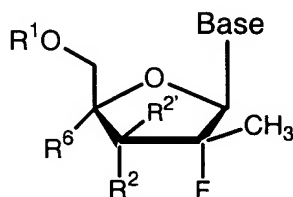
(a)



(b)

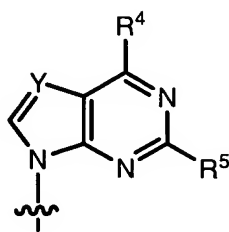
and wherein  $R^1$  is H,  $R^2$  is OH,  $R^{2'}$  is H,  $R^3$  is H, and  $R^4$  is  $NH_2$  or OH, and  $R^5$  is  $NH_2$ .

64. A method for the treatment or prophylaxis of a yellow fever virus infection comprising administering to a host an antivirally effective amount of a (2'R)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside ( $\beta$ -D or  $\beta$ -L) of the formula:

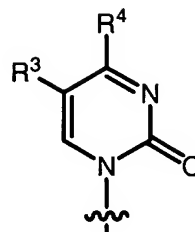


wherein

Base is selected from the group consisting of



(a)



(b)

Y is N or CH;

R<sup>1</sup> and R<sup>7</sup> are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid (or racemic mixture), a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein R<sup>1</sup> is H or phosphate; R<sup>2</sup> is H or phosphate; R<sup>1</sup> and R<sup>2</sup> or R<sup>7</sup> can also be linked with cyclic phosphate group;

R<sup>2</sup> and R<sup>2'</sup> are independently H, C<sub>1-4</sub> alkyl, C<sub>1-4</sub> alkenyl, C<sub>1-4</sub> alkynyl, vinyl, N<sub>3</sub>, CN, Cl, Br, F, I, NO<sub>2</sub>, C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkynyl), C(O)O(C<sub>1-4</sub> alkenyl), O(C<sub>1-4</sub> acyl), O(C<sub>1-4</sub> alkyl), O(C<sub>1-4</sub> alkenyl), S(C<sub>1-4</sub> acyl), S(C<sub>1-4</sub> alkyl), S(C<sub>1-4</sub> alkynyl), S(C<sub>1-4</sub> alkenyl), SO(C<sub>1-4</sub> acyl), SO(C<sub>1-4</sub> alkyl), SO(C<sub>1-4</sub> alkynyl), SO(C<sub>1-4</sub> alkenyl), SO<sub>2</sub>(C<sub>1-4</sub> acyl), SO<sub>2</sub>(C<sub>1-4</sub> alkyl), SO<sub>2</sub>(C<sub>1-4</sub> alkynyl), SO<sub>2</sub>(C<sub>1-4</sub> alkenyl), O<sub>3</sub>S(C<sub>1-4</sub> acyl), O<sub>3</sub>S(C<sub>1-4</sub> alkyl), O<sub>3</sub>S(C<sub>1-4</sub> alkenyl), NH<sub>2</sub>, NH(C<sub>1-4</sub> alkyl), NH(C<sub>1-4</sub> alkenyl), NH(C<sub>1-4</sub> alkynyl), NH(C<sub>1-4</sub> acyl), N(C<sub>1-4</sub> alkyl)<sub>2</sub>, N(C<sub>1-18</sub> acyl)<sub>2</sub>, wherein alkyl, alkynyl, alkenyl and vinyl are optionally substituted by N<sub>3</sub>, CN, one to three halogen (Cl, Br, F, I), NO<sub>2</sub>, C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkynyl), C(O)O(C<sub>1-4</sub> alkenyl), O(C<sub>1-4</sub> acyl), O(C<sub>1-4</sub> alkyl), O(C<sub>1-4</sub> alkenyl), S(C<sub>1-4</sub> acyl), S(C<sub>1-4</sub> alkyl), S(C<sub>1-4</sub> alkynyl), S(C<sub>1-4</sub> alkenyl), SO(C<sub>1-4</sub> acyl), SO(C<sub>1-4</sub> alkyl), SO(C<sub>1-4</sub> alkynyl), SO(C<sub>1-4</sub> alkenyl), SO<sub>2</sub>(C<sub>1-4</sub> acyl), SO<sub>2</sub>(C<sub>1-4</sub> alkyl), SO<sub>2</sub>(C<sub>1-4</sub> alkynyl), SO<sub>2</sub>(C<sub>1-4</sub> alkenyl), O<sub>3</sub>S(C<sub>1-4</sub> acyl), O<sub>3</sub>S(C<sub>1-4</sub> alkyl), O<sub>3</sub>S(C<sub>1-4</sub> alkenyl), NH<sub>2</sub>, NH(C<sub>1-4</sub> alkyl), NH(C<sub>1-4</sub>

alkenyl), NH(C<sub>1-4</sub> alkynyl), NH(C<sub>1-4</sub> acyl), N(C<sub>1-4</sub> alkyl)<sub>2</sub>, N(C<sub>1-4</sub> acyl)<sub>2</sub>, OR<sup>7</sup>; R<sup>2</sup> and R<sup>2'</sup> can be linked together to form a vinyl optionally substituted by one or two of N<sub>3</sub>, CN, Cl, Br, F, I, NO<sub>2</sub>;

5 R<sup>3</sup>, R<sup>4</sup> and R<sup>5</sup> are independently H, halogen including F, Cl, Br, I, OH, OR', SH, SR', NH<sub>2</sub>, NHR', NR'<sub>2</sub>, lower alkyl of C<sub>1</sub>-C<sub>6</sub>, halogenated (F, Cl, Br, I) lower alkyl of C<sub>1</sub>-C<sub>6</sub> such as CF<sub>3</sub> and CH<sub>2</sub>CH<sub>2</sub>F, lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CH<sub>2</sub>, halogenated (F, Cl, Br, I) lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CHCl, CH=CHBr and CH=CHI, lower alkynyl of C<sub>2</sub>-C<sub>6</sub> such as C≡CH, halogenated (F, Cl, Br, I) lower alkynyl of C<sub>2</sub>-C<sub>6</sub>,  
10 lower alkoxy of C<sub>1</sub>-C<sub>6</sub> such as CH<sub>2</sub>OH and CH<sub>2</sub>CH<sub>2</sub>OH, halogenated (F, Cl, Br, I) lower alkoxy of C<sub>1</sub>-C<sub>6</sub>, CO<sub>2</sub>H, CO<sub>2</sub>R', CONH<sub>2</sub>, CONHR', CONR'<sub>2</sub>, CH=CHCO<sub>2</sub>H, CH=CHCO<sub>2</sub>R';

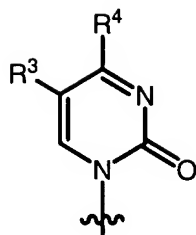
R' is an optionally substituted alkyl of C<sub>1</sub>-C<sub>12</sub> (particularly when the alkyl is an amino acid residue), cycloalkyl, optionally substituted alkynyl of C<sub>2</sub>-C<sub>6</sub>, optionally substituted lower alkenyl of C<sub>2</sub>-C<sub>6</sub>, or optionally substituted acyl;

R<sup>6</sup> is an optionally substituted alkyl (including lower alkyl), cyano (CN), CH<sub>3</sub>, OCH<sub>3</sub>, OCH<sub>2</sub>CH<sub>3</sub>, hydroxy methyl (CH<sub>2</sub>OH), fluoromethyl (CH<sub>2</sub>F), azido (N<sub>3</sub>), CHCN, CH<sub>2</sub>N<sub>3</sub>, CH<sub>2</sub>NH<sub>2</sub>, CH<sub>2</sub>NHCH<sub>3</sub>, CH<sub>2</sub>N(CH<sub>3</sub>)<sub>2</sub>,  
20 alkyne (optionally substituted), or fluoro;

or its pharmaceutically acceptable salt or prodrug thereof. optionally in a pharmaceutically acceptable carrier.

65. The method of claim 64, wherein

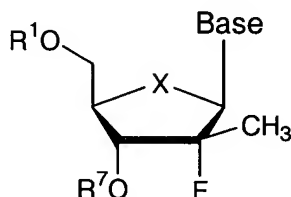
25 Base is



and  $R^1$  is H,  $R^2$  is OH,  $R^{2'}$  is H,  $R^3$  is H,  $R^4$  is  $\text{NH}_2$  or OH,  
and  $R^6$  is H.

5

66. A method for the treatment or prophylaxis of a yellow fever virus infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside ( $\beta$ -D or  $\beta$ -L) or its pharmaceutically acceptable salt or prodrug thereof of the structure:



10

wherein Base is a purine or pyrimidine base;

X is O, S,  $\text{CH}_2$ , Se, NH, N-alkyl, CHW (*R*, *S*, or racemic),  $\text{C(W)}_2$ , wherein W is F, Cl, Br, or I; and,

15

$R^1$  and  $R^7$  are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid, a

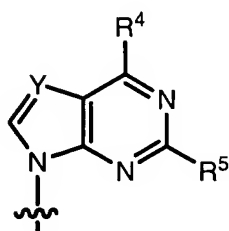
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carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein  $R^1$  or  $R^7$  is independently H or phosphate;  $R^1$  and  $R^7$  can also be linked with cyclic phosphate group and

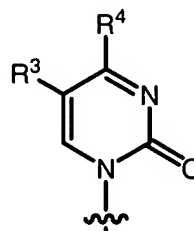
optionally in a pharmaceutically acceptable carrier.

67. The method of claim 66, wherein

Base is selected from the group consisting of:



(a)



(b)

Y is N or CH;

$R^3$ ,  $R^4$  and  $R^5$  are independently H, halogen including F, Cl, Br, I, OH, OR', SH, SR', NH<sub>2</sub>, NHR', NR'<sub>2</sub>, lower alkyl of C<sub>1</sub>-C<sub>6</sub>, halogenated (F, Cl, Br, I) lower alkyl of C<sub>1</sub>-C<sub>6</sub> such as CF<sub>3</sub> and CH<sub>2</sub>CH<sub>2</sub>F, lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CH<sub>2</sub>, halogenated (F, Cl, Br, I) lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CHCl, CH=CHBr and CH=CHI, lower alkynyl of C<sub>2</sub>-C<sub>6</sub> such as C≡CH, halogenated (F, Cl, Br, I) lower alkynyl of C<sub>2</sub>-C<sub>6</sub>, lower alkoxy of C<sub>1</sub>-C<sub>6</sub> such as CH<sub>2</sub>OH and CH<sub>2</sub>CH<sub>2</sub>OH, halogenated (F, Cl, Br, I) lower alkoxy of C<sub>1</sub>-C<sub>6</sub>, CO<sub>2</sub>H, CO<sub>2</sub>R', CONH<sub>2</sub>, CONHR', CONR'<sub>2</sub>, CH=CHCO<sub>2</sub>H, CH=CHCO<sub>2</sub>R'; and,

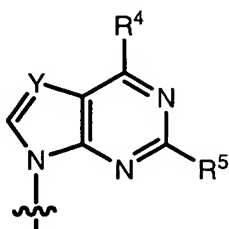
R' is an optionally substituted alkyl of C<sub>1</sub>-C<sub>12</sub> (particularly when the alkyl is an amino acid residue), cycloalkyl, optionally substituted alkynyl of C<sub>2</sub>-



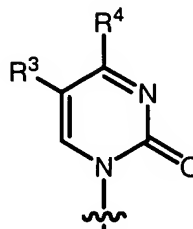
C<sub>6</sub>, optionally substituted lower alkenyl of C<sub>2</sub>-C<sub>6</sub>, or optionally substituted acyl.

68. The method of claim 66, wherein

5 Base is selected from the group consisting of (a) or (b):



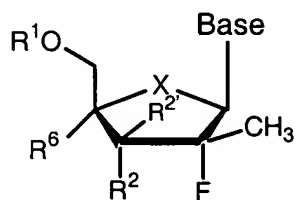
(a)



(b)

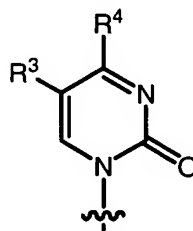
and wherein R<sup>1</sup> and R<sup>7</sup> are H, R<sup>3</sup> is H, and R<sup>4</sup> is NH<sub>2</sub> or OH, and R<sup>5</sup> is NH<sub>2</sub>.

10 69. A method for the treatment or prophylaxis of a yellow fever virus infection comprising administering to a host an antivirally effective amount of a (2'R)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside (β-D or β-L) of the formula:



wherein

15 Base is



X is O, S, CH<sub>2</sub>, Se, NH, N-alkyl, CHW (*R*, *S*, or racemic), C(W)<sub>2</sub>, wherein W is F, Cl, Br, or I;

R<sup>1</sup> and R<sup>7</sup> are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid (or racemic mixture), a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein R<sup>1</sup> is H or phosphate; R<sup>2</sup> is H or phosphate; R<sup>1</sup> and R<sup>2</sup> or R<sup>7</sup> can also be linked with cyclic phosphate group;

R<sup>2</sup> and R<sup>2'</sup> are independently H, C<sub>1-4</sub> alkyl, C<sub>1-4</sub> alkenyl, C<sub>1-4</sub> alkynyl, vinyl, N<sub>3</sub>, CN, Cl, Br, F, I, NO<sub>2</sub>, C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkynyl), C(O)O(C<sub>1-4</sub> alkenyl), O(C<sub>1-4</sub> acyl), O(C<sub>1-4</sub> alkyl), O(C<sub>1-4</sub> alkenyl), S(C<sub>1-4</sub> acyl), S(C<sub>1-4</sub> alkyl), S(C<sub>1-4</sub> alkynyl), S(C<sub>1-4</sub> alkenyl), SO(C<sub>1-4</sub> acyl), SO(C<sub>1-4</sub> alkyl), SO(C<sub>1-4</sub> alkynyl), SO(C<sub>1-4</sub> alkenyl), SO<sub>2</sub>(C<sub>1-4</sub> acyl), SO<sub>2</sub>(C<sub>1-4</sub> alkyl), SO<sub>2</sub>(C<sub>1-4</sub> alkynyl), SO<sub>2</sub>(C<sub>1-4</sub> alkenyl), O<sub>3</sub>S(C<sub>1-4</sub> acyl), O<sub>3</sub>S(C<sub>1-4</sub> alkyl), O<sub>3</sub>S(C<sub>1-4</sub> alkenyl), NH<sub>2</sub>, NH(C<sub>1-4</sub> alkyl), NH(C<sub>1-4</sub> alkenyl), NH(C<sub>1-4</sub> alkynyl), NH(C<sub>1-4</sub> acyl), N(C<sub>1-4</sub> alkyl)<sub>2</sub>, N(C<sub>1-18</sub> acyl)<sub>2</sub>, wherein alkyl, alkynyl, alkenyl and vinyl are optionally substituted by N<sub>3</sub>, CN, one to three halogen (Cl, Br, F, I), NO<sub>2</sub>, C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkynyl), C(O)O(C<sub>1-4</sub> alkenyl), O(C<sub>1-4</sub> acyl), O(C<sub>1-4</sub> alkyl), O(C<sub>1-4</sub> alkenyl), S(C<sub>1-4</sub> acyl), S(C<sub>1-4</sub> alkyl), S(C<sub>1-4</sub> alkynyl), S(C<sub>1-4</sub> alkenyl), SO(C<sub>1-4</sub> acyl), SO(C<sub>1-4</sub> alkyl), SO(C<sub>1-4</sub> alkynyl), SO(C<sub>1-4</sub> alkenyl), SO<sub>2</sub>(C<sub>1-4</sub> acyl), SO<sub>2</sub>(C<sub>1-4</sub> alkyl), SO<sub>2</sub>(C<sub>1-4</sub> alkynyl), SO<sub>2</sub>(C<sub>1-4</sub> alkenyl), O<sub>3</sub>S(C<sub>1-4</sub> acyl),

$O_3S(C_{1-4} \text{ alkyl})$ ,  $O_3S(C_{1-4} \text{ alkenyl})$ ,  $NH_2$ ,  $NH(C_{1-4} \text{ alkyl})$ ,  $NH(C_{1-4} \text{ alkenyl})$ ,  $NH(C_{1-4} \text{ alkynyl})$ ,  $NH(C_{1-4} \text{ acyl})$ ,  $N(C_{1-4} \text{ alkyl})_2$ ,  $N(C_{1-4} \text{ acyl})_2$ ,  $OR^7$ ;  $R^2$  and  $R^{2'}$  can be linked together to form a vinyl optionally substituted by one or two of  $N_3$ ,  $CN$ ,  $Cl$ ,  $Br$ ,  $F$ ,  $I$ ,  $NO_2$ ;

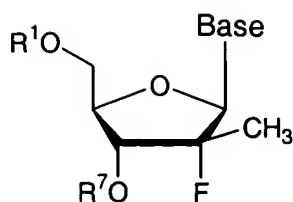
5  $R^3$  and  $R^4$  are independently  $H$ , halogen including  $F$ ,  $Cl$ ,  $Br$ ,  $I$ ,  $OH$ ,  $OR'$ ,  $SH$ ,  $SR'$ ,  $NH_2$ ,  $NHR'$ ,  $NR'_2$ , lower alkyl of  $C_1-C_6$ , halogenated ( $F$ ,  $Cl$ ,  $Br$ ,  $I$ ) lower alkyl of  $C_1-C_6$  such as  $CF_3$  and  $CH_2CH_2F$ , lower alkenyl of  $C_2-C_6$  such as  $CH=CH_2$ , halogenated ( $F$ ,  $Cl$ ,  $Br$ ,  $I$ ) lower alkenyl of  $C_2-C_6$  such as  $CH=CHCl$ ,  $CH=CHBr$  and  $CH=CHI$ , lower alkynyl of  $C_2-C_6$  such as  $C\equiv CH$ , halogenated ( $F$ ,  $Cl$ ,  $Br$ ,  $I$ ) lower alkynyl of  $C_2-C_6$ , lower  
10 alkoxy of  $C_1-C_6$  such as  $CH_2OH$  and  $CH_2CH_2OH$ , halogenated ( $F$ ,  $Cl$ ,  $Br$ ,  $I$ ) lower alkoxy of  $C_1-C_6$ ,  $CO_2H$ ,  $CO_2R'$ ,  $CONH_2$ ,  $CONHR'$ ,  $CONR'_2$ ,  $CH=CHCO_2H$ ,  $CH=CHCO_2R'$ ;

15  $R'$  is an optionally substituted alkyl of  $C_1-C_{12}$  (particularly when the alkyl is an amino acid residue), cycloalkyl, optionally substituted alkynyl of  $C_2-C_6$ , optionally substituted lower alkenyl of  $C_2-C_6$ , or optionally substituted acyl.

20  $R^6$  is an optionally substituted alkyl (including lower alkyl), cyano ( $CN$ ),  $CH_3$ ,  $OCH_3$ ,  $OCH_2CH_3$ , hydroxy methyl ( $CH_2OH$ ), fluoromethyl ( $CH_2F$ ), azido ( $N_3$ ),  $CHCN$ ,  $CH_2N_3$ ,  $CH_2NH_2$ ,  $CH_2NHCH_3$ ,  $CH_2N(CH_3)_2$ , alkyne (optionally substituted), or fluoro;

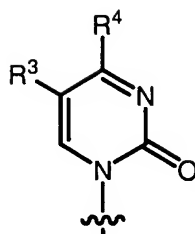
or its pharmaceutically acceptable salt or prodrug thereof, optionally in a pharmaceutically acceptable carrier.

25 70. A method for the treatment or prophylaxis of a yellow fever virus infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside ( $\beta$ -D or  $\beta$ -L) of the formula:



wherein

Base is



5

$R^1$  and  $R^7$  are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid, a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein  $R^1$  or  $R^7$  is independently H or phosphate;  $R^1$  and  $R^7$  can also be linked with cyclic phosphate group;

10

15

$R^3$  and  $R^4$  are independently H, halogen including F, Cl, Br, I, OH, OR', SH, SR', NH<sub>2</sub>, NHR', NR'<sub>2</sub>, lower alkyl of C<sub>1</sub>-C<sub>6</sub>, halogenated (F, Cl, Br, I) lower alkyl of C<sub>1</sub>-C<sub>6</sub> such as CF<sub>3</sub> and CH<sub>2</sub>CH<sub>2</sub>F, lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CH<sub>2</sub>, halogenated (F, Cl, Br, I) lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CHCl, CH=CHBr and CH=CHI, lower alkynyl of C<sub>2</sub>-C<sub>6</sub>

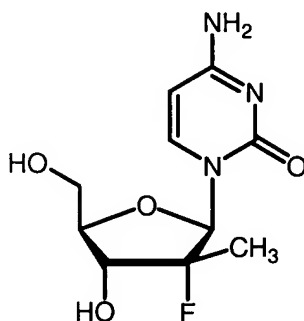
20

such as  $C\equiv CH$ , halogenated (F, Cl, Br, I) lower alkynyl of  $C_2-C_6$ , lower alkoxy of  $C_1-C_6$  such as  $CH_2OH$  and  $CH_2CH_2OH$ , halogenated (F, Cl, Br, I) lower alkoxy of  $C_1-C_6$ ,  $CO_2H$ ,  $CO_2R'$ ,  $CONH_2$ ,  $CONHR'$ ,  $CONR'_2$ ,  $CH=CHCO_2H$ ,  $CH=CHCO_2R'$ ;

5  $R'$  is an optionally substituted alkyl of  $C_1-C_{12}$  (particularly when the alkyl is an amino acid residue), cycloalkyl, optionally substituted alkynyl of  $C_2-C_6$ , optionally substituted lower alkenyl of  $C_2-C_6$ , or optionally substituted acyl;

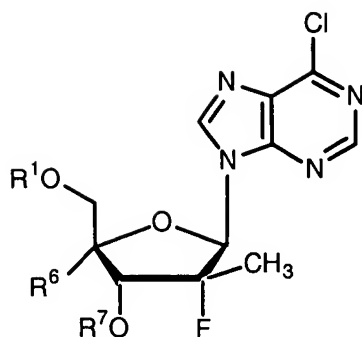
10 or its pharmaceutically acceptable salt or prodrug thereof, optionally in a pharmaceutically acceptable carrier.

15 71. A method for the treatment or prophylaxis of a yellow fever virus infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside ( $\beta$ -D) or its pharmaceutically acceptable salt or prodrug thereof of the formula:



optionally in a pharmaceutically acceptable carrier.

20 72. A method for the treatment or prophylaxis of a yellow fever virus infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside ( $\beta$ -D) or its pharmaceutically acceptable salt or prodrug thereof of the formula:



wherein

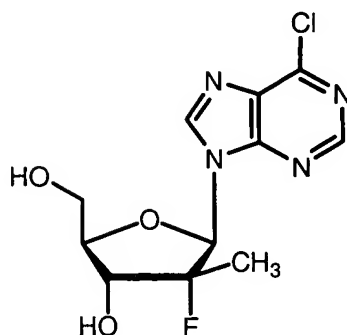
$R^1$  and  $R^7$  are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid, a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein  $R^1$  or  $R^7$  is independently H or phosphate;  $R^1$  and  $R^7$  can also be linked with cyclic phosphate group; and,

$R^6$  is an optionally substituted alkyl (including lower alkyl), cyano (CN),  $CH_3$ ,  $OCH_3$ ,  $OCH_2CH_3$ , hydroxy methyl ( $CH_2OH$ ), fluoromethyl ( $CH_2F$ ), azido ( $N_3$ ),  $CHCN$ ,  $CH_2N_3$ ,  $CH_2NH_2$ ,  $CH_2NHCH_3$ ,  $CH_2N(CH_3)_2$ , alkyne (optionally substituted), or fluoro, and

optionally in a pharmaceutically acceptable carrier.

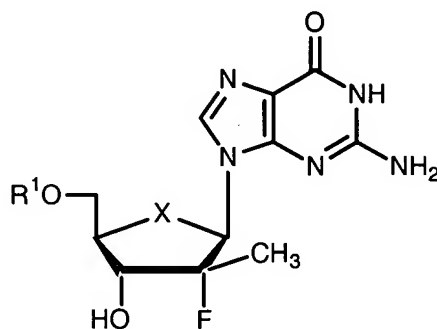
73. A method for the treatment or prophylaxis of a yellow fever virus infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-

fluoro-2'-C-methyl nucleoside ( $\beta$ -D) or its pharmaceutically acceptable salt or prodrug thereof of the formula:



optionally in a pharmaceutically acceptable carrier.

- 5                    74.     A method for the treatment or prophylaxis of a yellow fever virus infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside ( $\beta$ -D) or its pharmaceutically acceptable salt or prodrug thereof of the formula:



10                    wherein

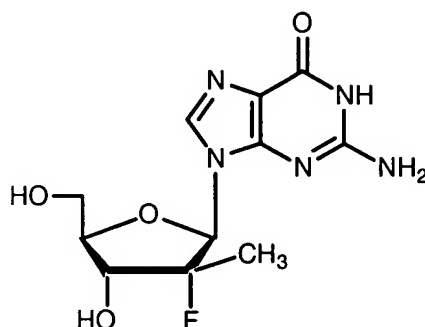
X is O, S, CH<sub>2</sub>, Se, NH, N-alkyl, CHW (*R*, *S*, or racemic), C(W)<sub>2</sub>, wherein W is F, Cl, Br, or I; and

15                    R<sup>1</sup> is H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl

sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid, a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein R<sup>1</sup> is H or phosphate;

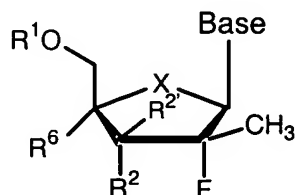
optionally in a pharmaceutically acceptable carrier

75. A method for the treatment or prophylaxis of a yellow fever virus infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside (β-D) or its pharmaceutically acceptable salt or prodrug thereof of the formula:



optionally in a pharmaceutically acceptable carrier.

76. A method for the treatment or prophylaxis of a West Nile virus infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside (β-D or β-L) of the formula:



wherein



Base is a purine or pyrimidine base;

X is O, S, CH<sub>2</sub>, Se, NH, N-alkyl, CHW (*R*, *S*, or racemic), C(W)<sub>2</sub>, wherein W is F, Cl, Br, or I;

R<sup>1</sup> and R<sup>7</sup> are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid (or racemic mixture), a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein R<sup>1</sup> is H or phosphate; R<sup>2</sup> is H or phosphate; R<sup>1</sup> and R<sup>2</sup> or R<sup>7</sup> can also be linked with cyclic phosphate group;

R<sup>2</sup> and R<sup>2'</sup> are independently H, C<sub>1-4</sub> alkyl, C<sub>1-4</sub> alkenyl, C<sub>1-4</sub> alkynyl, vinyl, N<sub>3</sub>, CN, Cl, Br, F, I, NO<sub>2</sub>, C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkynyl), C(O)O(C<sub>1-4</sub> alkenyl), O(C<sub>1-4</sub> acyl), O(C<sub>1-4</sub> alkyl), O(C<sub>1-4</sub> alkenyl), S(C<sub>1-4</sub> acyl), S(C<sub>1-4</sub> alkyl), S(C<sub>1-4</sub> alkynyl), S(C<sub>1-4</sub> alkenyl), SO(C<sub>1-4</sub> acyl), SO(C<sub>1-4</sub> alkyl), SO(C<sub>1-4</sub> alkynyl), SO(C<sub>1-4</sub> alkenyl), SO<sub>2</sub>(C<sub>1-4</sub> acyl), SO<sub>2</sub>(C<sub>1-4</sub> alkyl), SO<sub>2</sub>(C<sub>1-4</sub> alkynyl), SO<sub>2</sub>(C<sub>1-4</sub> alkenyl), O<sub>3</sub>S(C<sub>1-4</sub> acyl), O<sub>3</sub>S(C<sub>1-4</sub> alkyl), O<sub>3</sub>S(C<sub>1-4</sub> alkenyl), NH<sub>2</sub>, NH(C<sub>1-4</sub> alkyl), NH(C<sub>1-4</sub> alkenyl), NH(C<sub>1-4</sub> alkynyl), NH(C<sub>1-4</sub> acyl), N(C<sub>1-4</sub> alkyl)<sub>2</sub>, N(C<sub>1-18</sub> acyl)<sub>2</sub>, wherein alkyl, alkynyl, alkenyl and vinyl are optionally substituted by N<sub>3</sub>, CN, one to three halogen (Cl, Br, F, I), NO<sub>2</sub>, C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkynyl), C(O)O(C<sub>1-4</sub> alkenyl), O(C<sub>1-4</sub> acyl), O(C<sub>1-4</sub> alkyl), O(C<sub>1-4</sub> alkenyl), S(C<sub>1-4</sub> acyl), S(C<sub>1-4</sub> alkyl), S(C<sub>1-4</sub> alkynyl), S(C<sub>1-4</sub> alkenyl), SO(C<sub>1-4</sub> acyl), SO(C<sub>1-4</sub> alkyl), SO(C<sub>1-4</sub> alkynyl), SO(C<sub>1-4</sub> alkenyl), SO<sub>2</sub>(C<sub>1-4</sub> acyl),

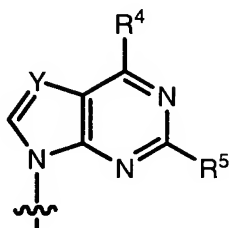
SO<sub>2</sub>(C<sub>1-4</sub> alkyl), SO<sub>2</sub>(C<sub>1-4</sub> alkynyl), SO<sub>2</sub>(C<sub>1-4</sub> alkenyl), O<sub>3</sub>S(C<sub>1-4</sub> acyl), O<sub>3</sub>S(C<sub>1-4</sub> alkyl), O<sub>3</sub>S(C<sub>1-4</sub> alkenyl), NH<sub>2</sub>, NH(C<sub>1-4</sub> alkyl), NH(C<sub>1-4</sub> alkenyl), NH(C<sub>1-4</sub> alkynyl), NH(C<sub>1-4</sub> acyl), N(C<sub>1-4</sub> alkyl)<sub>2</sub>, N(C<sub>1-4</sub> acyl)<sub>2</sub>, OR<sup>7</sup>; R<sup>2</sup> and R<sup>2'</sup> can be linked together to form a vinyl optionally substituted by one or two of N<sub>3</sub>, CN, Cl, Br, F, I, NO<sub>2</sub>;

R<sup>6</sup> is an optionally substituted alkyl (including lower alkyl), cyano (CN), CH<sub>3</sub>, OCH<sub>3</sub>, OCH<sub>2</sub>CH<sub>3</sub>, hydroxy methyl (CH<sub>2</sub>OH), fluoromethyl (CH<sub>2</sub>F), azido (N<sub>3</sub>), CHCN, CH<sub>2</sub>N<sub>3</sub>, CH<sub>2</sub>NH<sub>2</sub>, CH<sub>2</sub>NHCH<sub>3</sub>, CH<sub>2</sub>N(CH<sub>3</sub>)<sub>2</sub>, alkyne (optionally substituted), or fluoro;

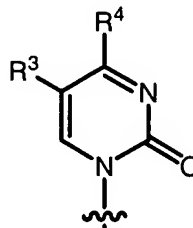
or its pharmaceutically acceptable salt or prodrug thereof, optionally in a pharmaceutically acceptable carrier.

77. The method of claim 76,

wherein Base is selected from the group consisting of:



(a)



(b)

Y is N or CH.

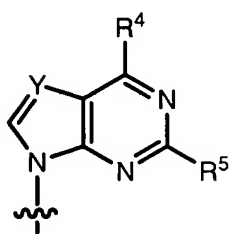
R<sup>3</sup>, R<sup>4</sup> and R<sup>5</sup> are independently H, halogen including F, Cl, Br, I, OH, OR', SH, SR', NH<sub>2</sub>, NHR', NR'<sub>2</sub>, lower alkyl of C<sub>1</sub>-C<sub>6</sub>, halogenated (F, Cl, Br, I) lower alkyl of C<sub>1</sub>-C<sub>6</sub> such as CF<sub>3</sub> and CH<sub>2</sub>CH<sub>2</sub>F, lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CH<sub>2</sub>, halogenated (F, Cl, Br, I) lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CHCl, CH=CHBr and CH=CHI, lower alkynyl of C<sub>2</sub>-C<sub>6</sub> such as C≡CH, halogenated (F, Cl, Br, I) lower alkynyl of C<sub>2</sub>-C<sub>6</sub>,

lower alkoxy of C<sub>1</sub>-C<sub>6</sub> such as CH<sub>2</sub>OH and CH<sub>2</sub>CH<sub>2</sub>OH, halogenated (F, Cl, Br, I) lower alkoxy of C<sub>1</sub>-C<sub>6</sub>, CO<sub>2</sub>H, CO<sub>2</sub>R', CONH<sub>2</sub>, CONHR', CONR'<sub>2</sub>, CH=CHCO<sub>2</sub>H, CH=CHCO<sub>2</sub>R'; and,

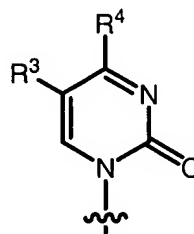
R' is an optionally substituted alkyl of C<sub>1</sub>-C<sub>12</sub> (particularly when the alkyl is an amino acid residue), cycloalkyl, optionally substituted alkynyl of C<sub>2</sub>-C<sub>6</sub>, optionally substituted lower alkenyl of C<sub>2</sub>-C<sub>6</sub>, or optionally substituted acyl.

78. The method of claim 76, wherein

Base is selected from the group consisting of (a) or (b):



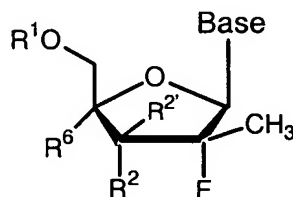
(a)



(b)

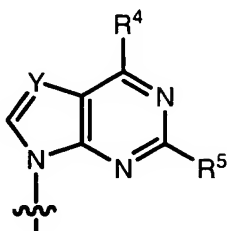
and wherein R<sup>1</sup> is H, R<sup>2</sup> is OH, R<sup>2'</sup> is H, R<sup>3</sup> is H, and R<sup>4</sup> is NH<sub>2</sub> or OH, and R<sup>5</sup> is NH<sub>2</sub>.

79. A method for the treatment or prophylaxis of a West Nile virus infection comprising administering to a host an antivirally effective amount of a (2'R)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside (β-D) of the formula:

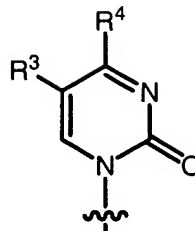


wherein

Base is selected from the group consisting of



(a)



(b)

5 Y is N or CH;

10  $R^1$  and  $R^7$  are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower  
15 alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid (or racemic mixture), a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein  $R^1$  is H or phosphate;  $R^2$  is H or phosphate;  $R^1$  and  $R^2$  or  $R^7$  can also be linked with cyclic phosphate group;

20  $R^2$  and  $R^{2'}$  are independently H,  $C_{1-4}$  alkyl,  $C_{1-4}$  alkenyl,  $C_{1-4}$  alkynyl, vinyl,  $N_3$ , CN, Cl, Br, F, I,  $NO_2$ ,  $C(O)O(C_{1-4}$  alkyl),  $C(O)O(C_{1-4}$  alkyl),  $C(O)O(C_{1-4}$  alkynyl),  $C(O)O(C_{1-4}$  alkenyl),  $O(C_{1-4}$  acyl),  $O(C_{1-4}$  alkyl),  $O(C_{1-4}$  alkenyl),  $S(C_{1-4}$  acyl),  $S(C_{1-4}$  alkyl),  $S(C_{1-4}$  alkynyl),  $S(C_{1-4}$  alkenyl),  $SO(C_{1-4}$  acyl),  $SO(C_{1-4}$  alkyl),  $SO(C_{1-4}$  alkynyl),  $SO(C_{1-4}$  alkenyl),  $SO_2(C_{1-4}$  acyl),  $SO_2(C_{1-4}$  alkyl),  $SO_2(C_{1-4}$  alkynyl),  $SO_2(C_{1-4}$  alkenyl),  
25  $O_3S(C_{1-4}$  acyl),  $O_3S(C_{1-4}$  alkyl),  $O_3S(C_{1-4}$  alkenyl),  $NH_2$ ,  $NH(C_{1-4}$

alkyl), NH(C<sub>1-4</sub> alkenyl), NH(C<sub>1-4</sub> alkynyl), NH(C<sub>1-4</sub> acyl), N(C<sub>1-4</sub> alkyl)<sub>2</sub>, N(C<sub>1-18</sub> acyl)<sub>2</sub>, wherein alkyl, alkynyl, alkenyl and vinyl are optionally substituted by N<sub>3</sub>, CN, one to three halogen (Cl, Br, F, I), NO<sub>2</sub>, C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkynyl), C(O)O(C<sub>1-4</sub> alkenyl), O(C<sub>1-4</sub> acyl), O(C<sub>1-4</sub> alkyl), O(C<sub>1-4</sub> alkenyl), S(C<sub>1-4</sub> acyl), S(C<sub>1-4</sub> alkyl), S(C<sub>1-4</sub> alkynyl), S(C<sub>1-4</sub> alkenyl), SO(C<sub>1-4</sub> acyl), SO(C<sub>1-4</sub> alkyl), SO(C<sub>1-4</sub> alkynyl), SO(C<sub>1-4</sub> alkenyl), SO<sub>2</sub>(C<sub>1-4</sub> acyl), SO<sub>2</sub>(C<sub>1-4</sub> alkyl), SO<sub>2</sub>(C<sub>1-4</sub> alkynyl), SO<sub>2</sub>(C<sub>1-4</sub> alkenyl), O<sub>3</sub>S(C<sub>1-4</sub> acyl), O<sub>3</sub>S(C<sub>1-4</sub> alkyl), O<sub>3</sub>S(C<sub>1-4</sub> alkenyl), NH<sub>2</sub>, NH(C<sub>1-4</sub> alkyl), NH(C<sub>1-4</sub> alkenyl), NH(C<sub>1-4</sub> alkynyl), NH(C<sub>1-4</sub> acyl), N(C<sub>1-4</sub> alkyl)<sub>2</sub>, N(C<sub>1-4</sub> acyl)<sub>2</sub>, OR<sup>7</sup>; R<sup>2</sup> and R<sup>2'</sup> can be linked together to form a vinyl optionally substituted by one or two of N<sub>3</sub>, CN, Cl, Br, F, I, NO<sub>2</sub>;

R<sup>3</sup>, R<sup>4</sup> and R<sup>5</sup> are independently H, halogen including F, Cl, Br, I, OH, OR', SH, SR', NH<sub>2</sub>, NHR', NR'<sub>2</sub>, lower alkyl of C<sub>1</sub>-C<sub>6</sub>, halogenated (F, Cl, Br, I) lower alkyl of C<sub>1</sub>-C<sub>6</sub> such as CF<sub>3</sub> and CH<sub>2</sub>CH<sub>2</sub>F, lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CH<sub>2</sub>, halogenated (F, Cl, Br, I) lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CHCl, CH=CHBr and CH=CHI, lower alkynyl of C<sub>2</sub>-C<sub>6</sub> such as C≡CH, halogenated (F, Cl, Br, I) lower alkynyl of C<sub>2</sub>-C<sub>6</sub>, lower alkoxy of C<sub>1</sub>-C<sub>6</sub> such as CH<sub>2</sub>OH and CH<sub>2</sub>CH<sub>2</sub>OH, halogenated (F, Cl, Br, I) lower alkoxy of C<sub>1</sub>-C<sub>6</sub>, CO<sub>2</sub>H, CO<sub>2</sub>R', CONH<sub>2</sub>, CONHR', CONR'<sub>2</sub>, CH=CHCO<sub>2</sub>H, CH=CHCO<sub>2</sub>R';

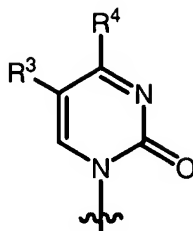
R' is an optionally substituted alkyl of C<sub>1</sub>-C<sub>12</sub> (particularly when the alkyl is an amino acid residue), cycloalkyl, optionally substituted alkynyl of C<sub>2</sub>-C<sub>6</sub>, optionally substituted lower alkenyl of C<sub>2</sub>-C<sub>6</sub>, or optionally substituted acyl;

R<sup>6</sup> is an optionally substituted alkyl (including lower alkyl), cyano (CN), CH<sub>3</sub>, OCH<sub>3</sub>, OCH<sub>2</sub>CH<sub>3</sub>, hydroxy methyl (CH<sub>2</sub>OH), fluoromethyl (CH<sub>2</sub>F), azido (N<sub>3</sub>), CHCN, CH<sub>2</sub>N<sub>3</sub>, CH<sub>2</sub>NH<sub>2</sub>, CH<sub>2</sub>NHCH<sub>3</sub>, CH<sub>2</sub>N(CH<sub>3</sub>)<sub>2</sub>, alkyne (optionally substituted), or fluoro;

or its pharmaceutically acceptable salt or prodrug thereof, optionally in a pharmaceutically acceptable carrier.

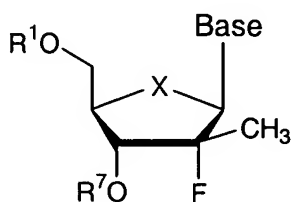
80. The method of claim 79, wherein

Base is



and  $R^1$  is H,  $R^2$  is OH,  $R^{2'}$  is H,  $R^3$  is H,  $R^4$  is  $\text{NH}_2$  or OH, and  $R^6$  is H.

81. A method for the treatment or prophylaxis of a West Nile virus infection comprising administering to a host an antivirally effective amount of a (2'R)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside ( $\beta$ -D or  $\beta$ -L) or its pharmaceutically acceptable salt or prodrug thereof of the structure:



wherein Base is a purine or pyrimidine base;

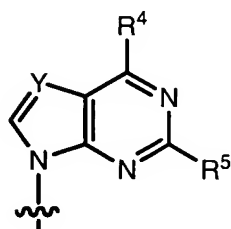
X is O, S,  $\text{CH}_2$ , Se, NH, N-alkyl, CHW (*R*, *S*, or racemic),  $\text{C(W)}_2$ , wherein W is F, Cl, Br, or I; and,

$R^1$  and  $R^7$  are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-

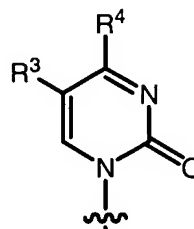
phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid, a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein R<sup>1</sup> or R<sup>7</sup> is independently H or phosphate; R<sup>1</sup> and R<sup>7</sup> can also be linked with cyclic phosphate group, and optionally a pharmaceutically acceptable carrier.

82. The method of claim 81, wherein

Base is selected from the group consisting of:



(a)



(b)

Y is N or CH;

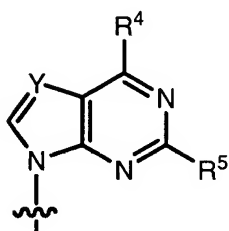
R<sup>3</sup>, R<sup>4</sup> and R<sup>5</sup> are independently H, halogen including F, Cl, Br, I, OH, OR', SH, SR', NH<sub>2</sub>, NHR', NR'<sub>2</sub>, lower alkyl of C<sub>1</sub>-C<sub>6</sub>, halogenated (F, Cl, Br, I) lower alkyl of C<sub>1</sub>-C<sub>6</sub> such as CF<sub>3</sub> and CH<sub>2</sub>CH<sub>2</sub>F, lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CH<sub>2</sub>, halogenated (F, Cl, Br, I) lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CHCl, CH=CHBr and CH=CHI, lower alkynyl of C<sub>2</sub>-C<sub>6</sub> such as C≡CH, halogenated (F, Cl, Br, I) lower alkynyl of C<sub>2</sub>-C<sub>6</sub>, lower alkoxy of C<sub>1</sub>-C<sub>6</sub> such as CH<sub>2</sub>OH and CH<sub>2</sub>CH<sub>2</sub>OH, halogenated

(F, Cl, Br, I) lower alkoxy of C<sub>1</sub>-C<sub>6</sub>, CO<sub>2</sub>H, CO<sub>2</sub>R', CONH<sub>2</sub>, CONHR', CONR'<sub>2</sub>, CH=CHCO<sub>2</sub>H, CH=CHCO<sub>2</sub>R'; and,

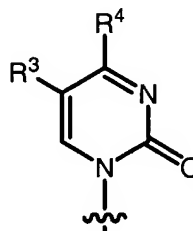
R' is an optionally substituted alkyl of C<sub>1</sub>-C<sub>12</sub> (particularly when the alkyl is an amino acid residue), cycloalkyl, optionally substituted alkynyl of C<sub>2</sub>-C<sub>6</sub>, optionally substituted lower alkenyl of C<sub>2</sub>-C<sub>6</sub>, or optionally substituted acyl.

83. The method of claim 81, wherein

Base is selected from the group consisting of (a) or (b):



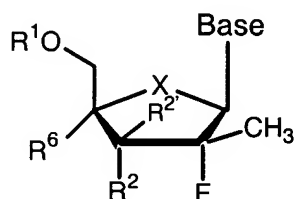
(a)



(b)

and wherein R<sup>1</sup> and R<sup>7</sup> are H, R<sup>3</sup> is H, and R<sup>4</sup> is NH<sub>2</sub> or OH, and R<sup>5</sup> is NH<sub>2</sub>.

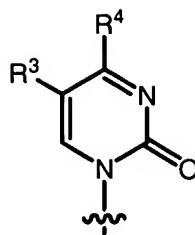
84. A method for the treatment or prophylaxis of a West Nile virus infection comprising administering to a host an antivirally effective amount of a (2'R)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside (β-D or β-L) of the formula:



wherein

Base is





X is O, S, CH<sub>2</sub>, Se, NH, N-alkyl, CHW (R, S, or racemic), C(W)<sub>2</sub>, wherein W is F, Cl, Br, or I;

R<sup>1</sup> and R<sup>7</sup> are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid (or racemic mixture), a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein R<sup>1</sup> is H or phosphate; R<sup>2</sup> is H or phosphate; R<sup>1</sup> and R<sup>2</sup> or R<sup>7</sup> can also be linked with cyclic phosphate group;

R<sup>2</sup> and R<sup>2'</sup> are independently H, C<sub>1-4</sub> alkyl, C<sub>1-4</sub> alkenyl, C<sub>1-4</sub> alkynyl, vinyl, N<sub>3</sub>, CN, Cl, Br, F, I, NO<sub>2</sub>, C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkynyl), C(O)O(C<sub>1-4</sub> alkenyl), O(C<sub>1-4</sub> acyl), O(C<sub>1-4</sub> alkyl), O(C<sub>1-4</sub> alkenyl), S(C<sub>1-4</sub> acyl), S(C<sub>1-4</sub> alkyl), S(C<sub>1-4</sub> alkynyl), S(C<sub>1-4</sub> alkenyl), SO(C<sub>1-4</sub> acyl), SO(C<sub>1-4</sub> alkyl), SO(C<sub>1-4</sub> alkynyl), SO(C<sub>1-4</sub> alkenyl), SO<sub>2</sub>(C<sub>1-4</sub> acyl), SO<sub>2</sub>(C<sub>1-4</sub> alkyl), SO<sub>2</sub>(C<sub>1-4</sub> alkynyl), SO<sub>2</sub>(C<sub>1-4</sub> alkenyl), O<sub>3</sub>S(C<sub>1-4</sub> acyl), O<sub>3</sub>S(C<sub>1-4</sub> alkyl), O<sub>3</sub>S(C<sub>1-4</sub> alkenyl), NH<sub>2</sub>, NH(C<sub>1-4</sub> alkyl), NH(C<sub>1-4</sub> alkenyl), NH(C<sub>1-4</sub> alkynyl), NH(C<sub>1-4</sub> acyl), N(C<sub>1-4</sub> alkyl)<sub>2</sub>, N(C<sub>1-18</sub> acyl)<sub>2</sub>, wherein alkyl, alkynyl, alkenyl and vinyl are optionally substituted by N<sub>3</sub>, CN, one to three halogen (Cl, Br, F, I),

5 NO<sub>2</sub>, C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkynyl),  
 C(O)O(C<sub>1-4</sub> alkenyl), O(C<sub>1-4</sub> acyl), O(C<sub>1-4</sub> alkyl), O(C<sub>1-4</sub> alkenyl), S(C<sub>1-4</sub>  
 4 acyl), S(C<sub>1-4</sub> alkyl), S(C<sub>1-4</sub> alkynyl), S(C<sub>1-4</sub> alkenyl), SO(C<sub>1-4</sub> acyl),  
 SO(C<sub>1-4</sub> alkyl), SO(C<sub>1-4</sub> alkynyl), SO(C<sub>1-4</sub> alkenyl), SO<sub>2</sub>(C<sub>1-4</sub> acyl),  
 SO<sub>2</sub>(C<sub>1-4</sub> alkyl), SO<sub>2</sub>(C<sub>1-4</sub> alkynyl), SO<sub>2</sub>(C<sub>1-4</sub> alkenyl), O<sub>3</sub>S(C<sub>1-4</sub> acyl),  
 O<sub>3</sub>S(C<sub>1-4</sub> alkyl), O<sub>3</sub>S(C<sub>1-4</sub> alkenyl), NH<sub>2</sub>, NH(C<sub>1-4</sub> alkyl), NH(C<sub>1-4</sub>  
 alkenyl), NH(C<sub>1-4</sub> alkynyl), NH(C<sub>1-4</sub> acyl), N(C<sub>1-4</sub> alkyl)<sub>2</sub>, N(C<sub>1-4</sub>  
 acyl)<sub>2</sub>, OR<sup>7</sup>; R<sup>2</sup> and R<sup>2'</sup> can be linked together to form a vinyl optionally  
 substituted by one or two of N<sub>3</sub>, CN, Cl, Br, F, I, NO<sub>2</sub>;

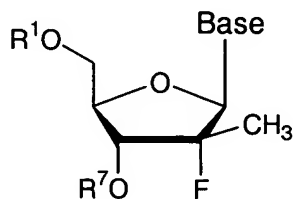
10 R<sup>3</sup> and R<sup>4</sup> are independently H, halogen including F, Cl, Br, I, OH, OR', SH,  
 SR', NH<sub>2</sub>, NHR', NR'<sub>2</sub>, lower alkyl of C<sub>1</sub>-C<sub>6</sub>, halogenated (F, Cl, Br, I)  
 lower alkyl of C<sub>1</sub>-C<sub>6</sub> such as CF<sub>3</sub> and CH<sub>2</sub>CH<sub>2</sub>F, lower alkenyl of C<sub>2</sub>-C<sub>6</sub>  
 such as CH=CH<sub>2</sub>, halogenated (F, Cl, Br, I) lower alkenyl of C<sub>2</sub>-C<sub>6</sub>  
 such as CH=CHCl, CH=CHBr and CH=CHI, lower alkynyl of C<sub>2</sub>-C<sub>6</sub>  
 15 such as C≡CH, halogenated (F, Cl, Br, I) lower alkynyl of C<sub>2</sub>-C<sub>6</sub>, lower  
 alkoxy of C<sub>1</sub>-C<sub>6</sub> such as CH<sub>2</sub>OH and CH<sub>2</sub>CH<sub>2</sub>OH, halogenated (F, Cl,  
 Br, I) lower alkoxy of C<sub>1</sub>-C<sub>6</sub>, CO<sub>2</sub>H, CO<sub>2</sub>R', CONH<sub>2</sub>, CONHR',  
 CONR'<sub>2</sub>, CH=CHCO<sub>2</sub>H, CH=CHCO<sub>2</sub>R'; and,

20 R' is an optionally substituted alkyl of C<sub>1</sub>-C<sub>12</sub> (particularly when the alkyl is an  
 amino acid residue), cycloalkyl, optionally substituted alkynyl of C<sub>2</sub>-  
 C<sub>6</sub>, optionally substituted lower alkenyl of C<sub>2</sub>-C<sub>6</sub>, or optionally  
 substituted acyl.

25 R<sup>6</sup> is an optionally substituted alkyl (including lower alkyl), cyano (CN), CH<sub>3</sub>,  
 OCH<sub>3</sub>, OCH<sub>2</sub>CH<sub>3</sub>, hydroxy methyl (CH<sub>2</sub>OH), fluoromethyl (CH<sub>2</sub>F),  
 azido (N<sub>3</sub>), CHCN, CH<sub>2</sub>N<sub>3</sub>, CH<sub>2</sub>NH<sub>2</sub>, CH<sub>2</sub>NHCH<sub>3</sub>, CH<sub>2</sub>N(CH<sub>3</sub>)<sub>2</sub>,  
 alkyne (optionally substituted), or fluoro;

or its pharmaceutically acceptable salt or prodrug thereof, optionally in a pharmaceutically  
 acceptable carrier.

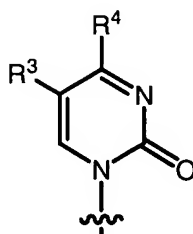
85. A method for the treatment or prophylaxis of a West Nile virus infection comprising administering to a host an antivirally effective amount of a (2'R)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside ( $\beta$ -D or  $\beta$ -L) of the formula:



5

wherein

Base is



10

$R^1$  and  $R^7$  are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid, a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein  $R^1$  or  $R^7$  is independently H or phosphate;  $R^1$  and  $R^7$  can also be linked with cyclic phosphate group;

15

20

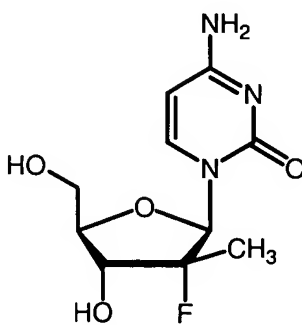
$R^3$  and  $R^4$  are independently H, halogen including F, Cl, Br, I, OH, OR', SH, SR', NH<sub>2</sub>, NHR', NR'<sub>2</sub>, lower alkyl of C<sub>1</sub>-C<sub>6</sub>, halogenated (F, Cl, Br, I)

lower alkyl of C<sub>1</sub>-C<sub>6</sub> such as CF<sub>3</sub> and CH<sub>2</sub>CH<sub>2</sub>F, lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CH<sub>2</sub>, halogenated (F, Cl, Br, I) lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CHCl, CH=CHBr and CH=CHI, lower alkynyl of C<sub>2</sub>-C<sub>6</sub> such as C≡CH, halogenated (F, Cl, Br, I) lower alkynyl of C<sub>2</sub>-C<sub>6</sub>, lower alkoxy of C<sub>1</sub>-C<sub>6</sub> such as CH<sub>2</sub>OH and CH<sub>2</sub>CH<sub>2</sub>OH, halogenated (F, Cl, Br, I) lower alkoxy of C<sub>1</sub>-C<sub>6</sub>, CO<sub>2</sub>H, CO<sub>2</sub>R', CONH<sub>2</sub>, CONHR', CONR'<sub>2</sub>, CH=CHCO<sub>2</sub>H, CH=CHCO<sub>2</sub>R';

R' is an optionally substituted alkyl of C<sub>1</sub>-C<sub>12</sub> (particularly when the alkyl is an amino acid residue), cycloalkyl, optionally substituted alkynyl of C<sub>2</sub>-C<sub>6</sub>, optionally substituted lower alkenyl of C<sub>2</sub>-C<sub>6</sub>, or optionally substituted acyl.

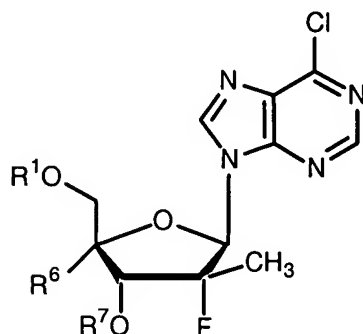
or its pharmaceutically acceptable salt or prodrug thereof, optionally in a pharmaceutically acceptable carrier.

86. A method for the treatment or prophylaxis of a West Nile virus infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside (β-D) or its pharmaceutically acceptable salt or prodrug thereof of the formula:



optionally in a pharmaceutically acceptable carrier.

87. A method for the treatment or prophylaxis of a West Nile virus infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside ( $\beta$ -D) or its pharmaceutically acceptable salt or prodrug thereof of the formula:



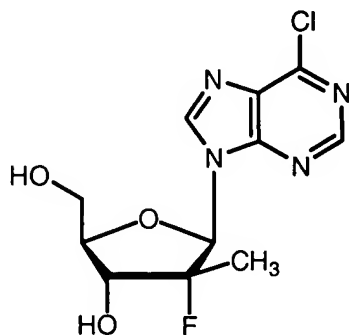
wherein

$R^1$  and  $R^7$  are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid, a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein  $R^1$  or  $R^7$  is independently H or phosphate;  $R^1$  and  $R^7$  can also be linked with cyclic phosphate group; and,

$R^6$  is an optionally substituted alkyl (including lower alkyl), cyano (CN),  $CH_3$ ,  $OCH_3$ ,  $OCH_2CH_3$ , hydroxy methyl ( $CH_2OH$ ), fluoromethyl ( $CH_2F$ ), azido ( $N_3$ ),  $CHCN$ ,  $CH_2N_3$ ,  $CH_2NH_2$ ,  $CH_2NHCH_3$ ,  $CH_2N(CH_3)_2$ , alkyne (optionally substituted), or fluoro and,

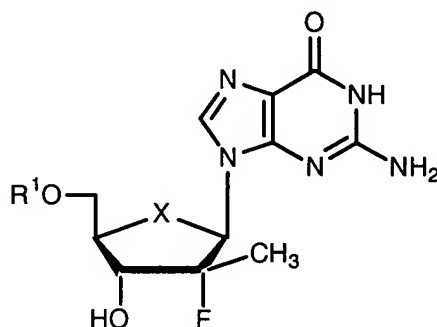
optionally in a pharmaceutically acceptable carrier.

88. A method for the treatment or prophylaxis of a West Nile virus infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside ( $\beta$ -D) or its pharmaceutically acceptable salt or prodrug thereof of the formula:



and optionally a pharmaceutically acceptable carrier.

89. A method for the treatment or prophylaxis of a West Nile virus infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside ( $\beta$ -D) or its pharmaceutically acceptable salt or prodrug thereof of the formula:



wherein

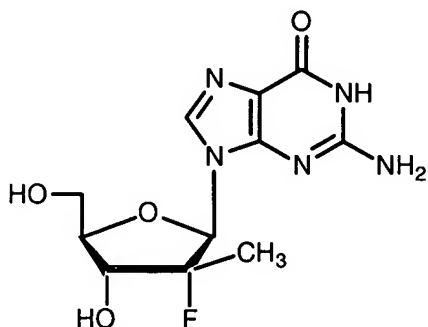
X is O, S, CH<sub>2</sub>, Se, NH, N-alkyl, CHW (*R*, *S*, or racemic), C(W)<sub>2</sub>, wherein W is F, Cl, Br, or I; and

R<sup>1</sup> is H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-

phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid, a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein R<sup>1</sup> is H or phosphate;

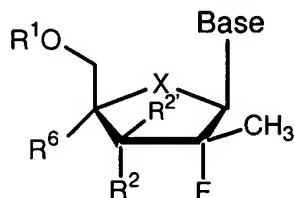
optionally in a pharmaceutically acceptable carrier.

90. A method for the treatment or prophylaxis of a West Nile virus infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside (β-*D*) or its pharmaceutically acceptable salt or prodrug thereof of the formula:



optionally in a pharmaceutically acceptable carrier.

91. A method for the treatment or prophylaxis of a Dengue virus infection comprising administering to a host an antivirally effective amount of a (2'R)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside ( $\beta$ -D or  $\beta$ -L) of the formula:



wherein

Base is a purine or pyrimidine base;

X is O, S, CH<sub>2</sub>, Se, NH, N-alkyl, CHW (*R*, *S*, or racemic), C(W)<sub>2</sub>, wherein W is F, Cl, Br, or I;

R<sup>1</sup> and R<sup>7</sup> are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid (or racemic mixture), a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein R<sup>1</sup> is H or phosphate; R<sup>2</sup> is H or phosphate; R<sup>1</sup> and R<sup>2</sup> or R<sup>7</sup> can also be linked with cyclic phosphate group;

R<sup>2</sup> and R<sup>2'</sup> are independently H, C<sub>1-4</sub> alkyl, C<sub>1-4</sub> alkenyl, C<sub>1-4</sub> alkynyl, vinyl, N<sub>3</sub>, CN, Cl, Br, F, I, NO<sub>2</sub>, C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkenyl), C(O)O(C<sub>1-4</sub> alkynyl), C(O)O(C<sub>1-4</sub> alkenyl), O(C<sub>1-4</sub> acyl), O(C<sub>1-4</sub> alkyl), O(C<sub>1-4</sub> alkenyl), S(C<sub>1-4</sub> acyl), S(C<sub>1-4</sub> alkyl), S(C<sub>1-4</sub> alkynyl), S(C<sub>1-4</sub> alkenyl), SO(C<sub>1-4</sub> acyl), SO(C<sub>1-4</sub> alkyl), SO(C<sub>1-4</sub> alkynyl), SO(C<sub>1-4</sub> alkenyl),



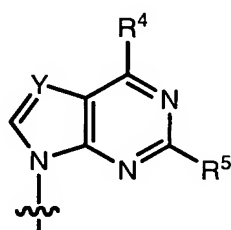
5 SO<sub>2</sub>(C<sub>1-4</sub> acyl), SO<sub>2</sub>(C<sub>1-4</sub> alkyl), SO<sub>2</sub>(C<sub>1-4</sub> alkynyl), SO<sub>2</sub>(C<sub>1-4</sub> alkenyl),  
 O<sub>3</sub>S(C<sub>1-4</sub> acyl), O<sub>3</sub>S(C<sub>1-4</sub> alkyl), O<sub>3</sub>S(C<sub>1-4</sub> alkenyl), NH<sub>2</sub>, NH(C<sub>1-4</sub>  
 alkyl), NH(C<sub>1-4</sub> alkenyl), NH(C<sub>1-4</sub> alkynyl), NH(C<sub>1-4</sub> acyl), N(C<sub>1-4</sub>  
 alkyl)<sub>2</sub>, N(C<sub>1-18</sub> acyl)<sub>2</sub>, wherein alkyl, alkynyl, alkenyl and vinyl are  
 10 optionally substituted by N<sub>3</sub>, CN, one to three halogen (Cl, Br, F, I),  
 NO<sub>2</sub>, C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkyl), C(O)O(C<sub>1-4</sub> alkynyl),  
 C(O)O(C<sub>1-4</sub> alkenyl), O(C<sub>1-4</sub> acyl), O(C<sub>1-4</sub> alkyl), O(C<sub>1-4</sub> alkenyl), S(C<sub>1-4</sub>  
 acyl), S(C<sub>1-4</sub> alkyl), S(C<sub>1-4</sub> alkynyl), S(C<sub>1-4</sub> alkenyl), SO(C<sub>1-4</sub> acyl),  
 SO(C<sub>1-4</sub> alkyl), SO(C<sub>1-4</sub> alkynyl), SO(C<sub>1-4</sub> alkenyl), SO<sub>2</sub>(C<sub>1-4</sub> acyl),  
 15 SO<sub>2</sub>(C<sub>1-4</sub> alkyl), SO<sub>2</sub>(C<sub>1-4</sub> alkynyl), SO<sub>2</sub>(C<sub>1-4</sub> alkenyl), O<sub>3</sub>S(C<sub>1-4</sub> acyl),  
 O<sub>3</sub>S(C<sub>1-4</sub> alkyl), O<sub>3</sub>S(C<sub>1-4</sub> alkenyl), NH<sub>2</sub>, NH(C<sub>1-4</sub> alkyl), NH(C<sub>1-4</sub>  
 alkenyl), NH(C<sub>1-4</sub> alkynyl), NH(C<sub>1-4</sub> acyl), N(C<sub>1-4</sub> alkyl)<sub>2</sub>, N(C<sub>1-4</sub>  
 acyl)<sub>2</sub>, OR<sup>7</sup>; R<sup>2</sup> and R<sup>2'</sup> can be linked together to form a vinyl optionally  
 substituted by one or two of N<sub>3</sub>, CN, Cl, Br, F, I, NO<sub>2</sub>;

15 R<sup>6</sup> is an optionally substituted alkyl (including lower alkyl), cyano (CN), CH<sub>3</sub>,  
 OCH<sub>3</sub>, OCH<sub>2</sub>CH<sub>3</sub>, hydroxy methyl (CH<sub>2</sub>OH), fluoromethyl (CH<sub>2</sub>F),  
 azido (N<sub>3</sub>), CHCN, CH<sub>2</sub>N<sub>3</sub>, CH<sub>2</sub>NH<sub>2</sub>, CH<sub>2</sub>NHCH<sub>3</sub>, CH<sub>2</sub>N(CH<sub>3</sub>)<sub>2</sub>,  
 alkyne (optionally substituted), or fluoro;

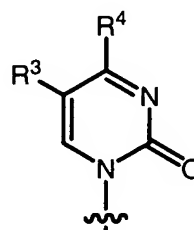
20 or its pharmaceutically acceptable salt or prodrug thereof, optionally in a pharmaceutically  
 acceptable carrier.

92. The method of claim 91,

wherein Base is selected from the group consisting of:



(a)



(b)

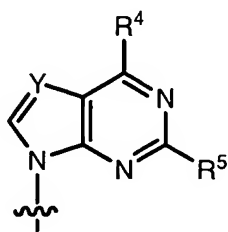
Y is N or CH.

$R^3$ ,  $R^4$  and  $R^5$  are independently H, halogen including F, Cl, Br, I, OH, OR', SH, SR', NH<sub>2</sub>, NHR', NR'<sub>2</sub>, lower alkyl of C<sub>1</sub>-C<sub>6</sub>, halogenated (F, Cl, Br, I) lower alkyl of C<sub>1</sub>-C<sub>6</sub> such as CF<sub>3</sub> and CH<sub>2</sub>CH<sub>2</sub>F, lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CH<sub>2</sub>, halogenated (F, Cl, Br, I) lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CHCl, CH=CHBr and CH=CHI, lower alkynyl of C<sub>2</sub>-C<sub>6</sub> such as C≡CH, halogenated (F, Cl, Br, I) lower alkynyl of C<sub>2</sub>-C<sub>6</sub>, lower alkoxy of C<sub>1</sub>-C<sub>6</sub> such as CH<sub>2</sub>OH and CH<sub>2</sub>CH<sub>2</sub>OH, halogenated (F, Cl, Br, I) lower alkoxy of C<sub>1</sub>-C<sub>6</sub>, CO<sub>2</sub>H, CO<sub>2</sub>R', CONH<sub>2</sub>, CONHR', CONR'<sub>2</sub>, CH=CHCO<sub>2</sub>H, CH=CHCO<sub>2</sub>R'; and,

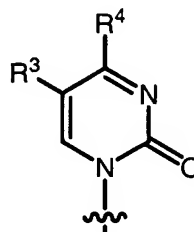
R' is an optionally substituted alkyl of C<sub>1</sub>-C<sub>12</sub> (particularly when the alkyl is an amino acid residue), cycloalkyl, optionally substituted alkynyl of C<sub>2</sub>-C<sub>6</sub>, optionally substituted lower alkenyl of C<sub>2</sub>-C<sub>6</sub>, or optionally substituted acyl.

93. The method of claim 91, wherein

Base is selected from the group consisting of (a) or (b):



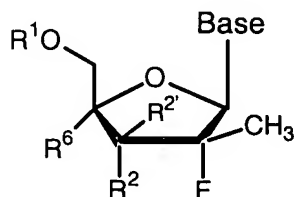
(a)



(b)

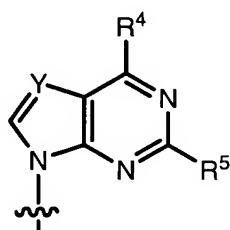
and wherein R<sup>1</sup> is H, R<sup>2</sup> is OH, R<sup>2'</sup> is H, R<sup>3</sup> is H, and R<sup>4</sup> is NH<sub>2</sub> or OH, and R<sup>5</sup> is NH<sub>2</sub>.

94. A method for the treatment or prophylaxis of a Dengue virus infection comprising administering to a host an antivirally effective amount of a (2'R)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside ( $\beta$ -D) of the formula:

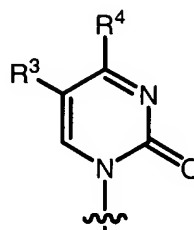


wherein

Base is selected from the group consisting of



(a)



(b)

Y is N or CH;

$R^1$  and  $R^7$  are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid (or racemic mixture), a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein  $R^1$  is H or phosphate;

$R^2$  is H or phosphate;  $R^1$  and  $R^2$  or  $R^7$  can also be linked with cyclic phosphate group;

$R^2$  and  $R^{2'}$  are independently H,  $C_{1-4}$  alkyl,  $C_{1-4}$  alkenyl,  $C_{1-4}$  alkynyl, vinyl,  $N_3$ , CN, Cl, Br, F, I,  $NO_2$ ,  $C(O)O(C_{1-4}$  alkyl),  $C(O)O(C_{1-4}$  alkyl),  $C(O)O(C_{1-4}$  alkynyl),  $C(O)O(C_{1-4}$  alkenyl),  $O(C_{1-4}$  acyl),  $O(C_{1-4}$  alkyl),  $O(C_{1-4}$  alkenyl),  $S(C_{1-4}$  acyl),  $S(C_{1-4}$  alkyl),  $S(C_{1-4}$  alkynyl),  $S(C_{1-4}$  alkenyl),  $SO(C_{1-4}$  acyl),  $SO(C_{1-4}$  alkyl),  $SO(C_{1-4}$  alkynyl),  $SO(C_{1-4}$  alkenyl),  $SO_2(C_{1-4}$  acyl),  $SO_2(C_{1-4}$  alkyl),  $SO_2(C_{1-4}$  alkynyl),  $SO_2(C_{1-4}$  alkenyl),  $O_3S(C_{1-4}$  acyl),  $O_3S(C_{1-4}$  alkyl),  $O_3S(C_{1-4}$  alkenyl),  $NH_2$ ,  $NH(C_{1-4}$  alkyl),  $NH(C_{1-4}$  alkenyl),  $NH(C_{1-4}$  alkynyl),  $NH(C_{1-4}$  acyl),  $N(C_{1-4}$  alkyl)<sub>2</sub>,  $N(C_{1-18}$  acyl)<sub>2</sub>, wherein alkyl, alkynyl, alkenyl and vinyl are optionally substituted by  $N_3$ , CN, one to three halogen (Cl, Br, F, I),  $NO_2$ ,  $C(O)O(C_{1-4}$  alkyl),  $C(O)O(C_{1-4}$  alkyl),  $C(O)O(C_{1-4}$  alkynyl),  $C(O)O(C_{1-4}$  alkenyl),  $O(C_{1-4}$  acyl),  $O(C_{1-4}$  alkyl),  $O(C_{1-4}$  alkenyl),  $S(C_{1-4}$  acyl),  $S(C_{1-4}$  alkyl),  $S(C_{1-4}$  alkynyl),  $S(C_{1-4}$  alkenyl),  $SO(C_{1-4}$  acyl),  $SO(C_{1-4}$  alkyl),  $SO(C_{1-4}$  alkynyl),  $SO(C_{1-4}$  alkenyl),  $SO_2(C_{1-4}$  acyl),  $SO_2(C_{1-4}$  alkyl),  $SO_2(C_{1-4}$  alkynyl),  $SO_2(C_{1-4}$  alkenyl),  $O_3S(C_{1-4}$  acyl),  $O_3S(C_{1-4}$  alkyl),  $O_3S(C_{1-4}$  alkenyl),  $NH_2$ ,  $NH(C_{1-4}$  alkyl),  $NH(C_{1-4}$  alkenyl),  $NH(C_{1-4}$  alkynyl),  $NH(C_{1-4}$  acyl),  $N(C_{1-4}$  alkyl)<sub>2</sub>,  $N(C_{1-4}$  acyl)<sub>2</sub>,  $OR^7$ ;  $R^2$  and  $R^{2'}$  can be linked together to form a vinyl optionally substituted by one or two of  $N_3$ , CN, Cl, Br, F, I,  $NO_2$ ;

$R^3$ ,  $R^4$  and  $R^5$  are independently H, halogen including F, Cl, Br, I, OH,  $OR'$ , SH,  $SR'$ ,  $NH_2$ ,  $NHR'$ ,  $NR'_2$ , lower alkyl of  $C_1-C_6$ , halogenated (F, Cl, Br, I) lower alkyl of  $C_1-C_6$  such as  $CF_3$  and  $CH_2CH_2F$ , lower alkenyl of  $C_2-C_6$  such as  $CH=CH_2$ , halogenated (F, Cl, Br, I) lower alkenyl of  $C_2-C_6$  such as  $CH=CHCl$ ,  $CH=CHBr$  and  $CH=CHI$ , lower alkynyl of  $C_2-C_6$  such as  $C\equiv CH$ , halogenated (F, Cl, Br, I) lower alkynyl of  $C_2-C_6$ , lower alkoxy of  $C_1-C_6$  such as  $CH_2OH$  and  $CH_2CH_2OH$ , halogenated (F, Cl, Br, I) lower alkoxy of  $C_1-C_6$ ,  $CO_2H$ ,  $CO_2R'$ ,  $CONH_2$ ,  $CONHR'$ ,  $CONR'_2$ ,  $CH=CHCO_2H$ ,  $CH=CHCO_2R'$ ;

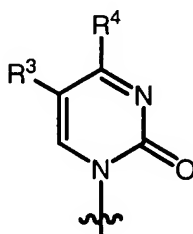
R' is an optionally substituted alkyl of C<sub>1</sub>-C<sub>12</sub> (particularly when the alkyl is an amino acid residue), cycloalkyl, optionally substituted alkynyl of C<sub>2</sub>-C<sub>6</sub>, optionally substituted lower alkenyl of C<sub>2</sub>-C<sub>6</sub>, or optionally substituted acyl;

5 R<sup>6</sup> is an optionally substituted alkyl (including lower alkyl), cyano (CN), CH<sub>3</sub>, OCH<sub>3</sub>, OCH<sub>2</sub>CH<sub>3</sub>, hydroxy methyl (CH<sub>2</sub>OH), fluoromethyl (CH<sub>2</sub>F), azido (N<sub>3</sub>), CHCN, CH<sub>2</sub>N<sub>3</sub>, CH<sub>2</sub>NH<sub>2</sub>, CH<sub>2</sub>NHCH<sub>3</sub>, CH<sub>2</sub>N(CH<sub>3</sub>)<sub>2</sub>, alkyne (optionally substituted), or fluoro;

10 or its pharmaceutically acceptable salt or prodrug thereof, optionally in a pharmaceutically acceptable carrier.

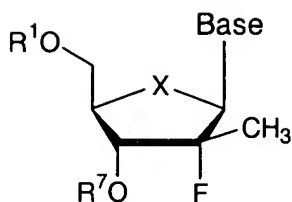
95. The method of claim 94, wherein

Base is



15 and R<sup>1</sup> is H, R<sup>2</sup> is OH, R<sup>2'</sup> is H, R<sup>3</sup> is H, R<sup>4</sup> is NH<sub>2</sub> or OH, and R<sup>6</sup> is H.

20 96. A method for the treatment or prophylaxis of a Dengue virus infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside (β-D or β-L) or its pharmaceutically acceptable salt or prodrug thereof of the structure:



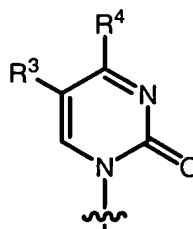
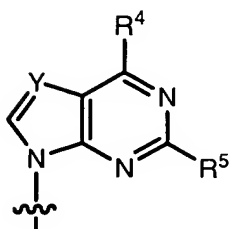
wherein Base is a purine or pyrimidine base;

X is O, S, CH<sub>2</sub>, Se, NH, N-alkyl, CHW (*R*, *S*, or racemic), C(W)<sub>2</sub>, wherein W is F, Cl, Br, or I; and,

5           R<sup>1</sup> and R<sup>7</sup> are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, 10           sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid, a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of 15           providing a compound wherein R<sup>1</sup> or R<sup>7</sup> is independently H or phosphate; R<sup>1</sup> and R<sup>7</sup> can also be linked with cyclic phosphate group, and optionally a pharmaceutically acceptable carrier.

97.   The method of claim 96, wherein

20           Base is selected from the group consisting of:



(a)

(b)

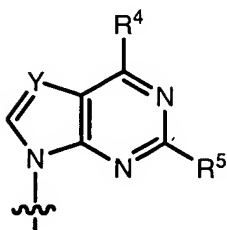
Y is N or CH;

$R^3$ ,  $R^4$  and  $R^5$  are independently H, halogen including F, Cl, Br, I, OH, OR', SH, SR', NH<sub>2</sub>, NHR', NR'<sub>2</sub>, lower alkyl of C<sub>1</sub>-C<sub>6</sub>, halogenated (F, Cl, Br, I) lower alkyl of C<sub>1</sub>-C<sub>6</sub> such as CF<sub>3</sub> and CH<sub>2</sub>CH<sub>2</sub>F, lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CH<sub>2</sub>, halogenated (F, Cl, Br, I) lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CHCl, CH=CHBr and CH=CHI, lower alkynyl of C<sub>2</sub>-C<sub>6</sub> such as C≡CH, halogenated (F, Cl, Br, I) lower alkynyl of C<sub>2</sub>-C<sub>6</sub>, lower alkoxy of C<sub>1</sub>-C<sub>6</sub> such as CH<sub>2</sub>OH and CH<sub>2</sub>CH<sub>2</sub>OH, halogenated (F, Cl, Br, I) lower alkoxy of C<sub>1</sub>-C<sub>6</sub>, CO<sub>2</sub>H, CO<sub>2</sub>R', CONH<sub>2</sub>, CONHR', CONR'<sub>2</sub>, CH=CHCO<sub>2</sub>H, CH=CHCO<sub>2</sub>R'; and,

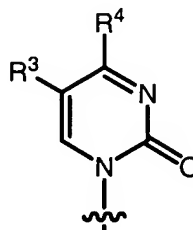
R' is an optionally substituted alkyl of C<sub>1</sub>-C<sub>12</sub> (particularly when the alkyl is an amino acid residue), cycloalkyl, optionally substituted alkynyl of C<sub>2</sub>-C<sub>6</sub>, optionally substituted lower alkenyl of C<sub>2</sub>-C<sub>6</sub>, or optionally substituted acyl.

98. The method of claim 96, wherein

Base is selected from the group consisting of (a) or (b):



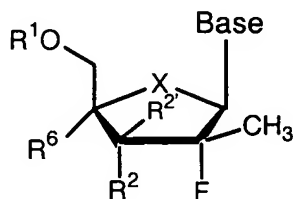
(a)



(b)

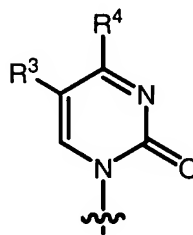
and wherein R<sup>1</sup> and R<sup>7</sup> are H, R<sup>3</sup> is H, and R<sup>4</sup> is NH<sub>2</sub> or OH, and R<sup>5</sup> is NH<sub>2</sub>.

99. A method for the treatment or prophylaxis of a Dengue virus infection comprising administering to a host an antivirally effective amount of a (2'R)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside ( $\beta$ -D or  $\beta$ -L) of the formula:



5 wherein

Base is



X is O, S, CH<sub>2</sub>, Se, NH, N-alkyl, CHW (*R*, *S*, or racemic), C(W)<sub>2</sub>, wherein W is F, Cl, Br, or I;

10 R<sup>1</sup> and R<sup>7</sup> are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including  
15 methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid (or racemic mixture), a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein R<sup>1</sup> is H or phosphate; R<sup>2</sup> is H or phosphate; R<sup>1</sup> and R<sup>2</sup> or R<sup>7</sup> can also be linked with cyclic phosphate group;  
20



$R^2$  and  $R^{2'}$  are independently H,  $C_{1-4}$  alkyl,  $C_{1-4}$  alkenyl,  $C_{1-4}$  alkynyl, vinyl,  $N_3$ ,  
 CN, Cl, Br, F, I,  $NO_2$ ,  $C(O)O(C_{1-4} \text{ alkyl})$ ,  $C(O)O(C_{1-4} \text{ alkenyl})$ ,  $C(O)O(C_{1-4} \text{ alkynyl})$ ,  
 5  $C(O)O(C_{1-4} \text{ acyl})$ ,  $O(C_{1-4} \text{ alkyl})$ ,  $O(C_{1-4} \text{ alkenyl})$ ,  $S(C_{1-4} \text{ alkyl})$ ,  $S(C_{1-4} \text{ alkenyl})$ ,  
 $SO(C_{1-4} \text{ alkyl})$ ,  $SO(C_{1-4} \text{ alkenyl})$ ,  $SO_2(C_{1-4} \text{ alkyl})$ ,  $SO_2(C_{1-4} \text{ alkenyl})$ ,  
 $O_3S(C_{1-4} \text{ alkyl})$ ,  $O_3S(C_{1-4} \text{ alkenyl})$ ,  $NH_2$ ,  $NH(C_{1-4} \text{ alkyl})$ ,  $NH(C_{1-4} \text{ alkenyl})$ ,  
 10  $NH(C_{1-4} \text{ alkynyl})$ ,  $N(C_{1-4} \text{ alkyl})_2$ ,  $N(C_{1-18} \text{ acyl})_2$ , wherein alkyl, alkynyl, alkenyl and vinyl are  
 optionally substituted by  $N_3$ , CN, one to three halogen (Cl, Br, F, I),  
 $NO_2$ ,  $C(O)O(C_{1-4} \text{ alkyl})$ ,  $C(O)O(C_{1-4} \text{ alkenyl})$ ,  $C(O)O(C_{1-4} \text{ alkynyl})$ ,  
 $C(O)O(C_{1-4} \text{ acyl})$ ,  $O(C_{1-4} \text{ alkyl})$ ,  $O(C_{1-4} \text{ alkenyl})$ ,  $S(C_{1-4} \text{ alkyl})$ ,  
 15  $S(C_{1-4} \text{ alkenyl})$ ,  $SO(C_{1-4} \text{ alkyl})$ ,  $SO(C_{1-4} \text{ alkenyl})$ ,  $SO_2(C_{1-4} \text{ alkyl})$ ,  
 $SO_2(C_{1-4} \text{ alkenyl})$ ,  $O_3S(C_{1-4} \text{ alkyl})$ ,  $O_3S(C_{1-4} \text{ alkenyl})$ ,  $NH_2$ ,  $NH(C_{1-4} \text{ alkyl})$ ,  
 $NH(C_{1-4} \text{ alkenyl})$ ,  $NH(C_{1-4} \text{ alkynyl})$ ,  $N(C_{1-4} \text{ alkyl})_2$ ,  $N(C_{1-4} \text{ acyl})_2$ ,  $OR^7$ ;  $R^2$  and  $R^{2'}$  can be linked together to form a vinyl optionally  
 substituted by one or two of  $N_3$ , CN, Cl, Br, F, I,  $NO_2$ ;

20  $R^3$  and  $R^4$  are independently H, halogen including F, Cl, Br, I, OH,  $OR'$ , SH,  
 $SR'$ ,  $NH_2$ ,  $NHR'$ ,  $NR'_2$ , lower alkyl of  $C_1-C_6$ , halogenated (F, Cl, Br, I)  
 lower alkyl of  $C_1-C_6$  such as  $CF_3$  and  $CH_2CH_2F$ , lower alkenyl of  $C_2-C_6$   
 such as  $CH=CH_2$ , halogenated (F, Cl, Br, I) lower alkenyl of  $C_2-C_6$   
 such as  $CH=CHCl$ ,  $CH=CHBr$  and  $CH=CHI$ , lower alkynyl of  $C_2-C_6$   
 25 such as  $C\equiv CH$ , halogenated (F, Cl, Br, I) lower alkynyl of  $C_2-C_6$ , lower  
 alkoxy of  $C_1-C_6$  such as  $CH_2OH$  and  $CH_2CH_2OH$ , halogenated (F, Cl,  
 Br, I) lower alkoxy of  $C_1-C_6$ ,  $CO_2H$ ,  $CO_2R'$ ,  $CONH_2$ ,  $CONHR'$ ,  
 $CONR'_2$ ,  $CH=CHCO_2H$ ,  $CH=CHCO_2R'$ ; and,

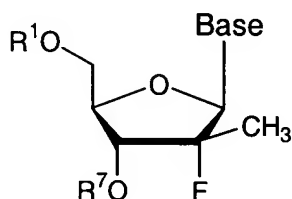
30  $R'$  is an optionally substituted alkyl of  $C_1-C_{12}$  (particularly when the alkyl is an  
 amino acid residue), cycloalkyl, optionally substituted alkynyl of  $C_2-$

C<sub>6</sub>, optionally substituted lower alkenyl of C<sub>2</sub>-C<sub>6</sub>, or optionally substituted acyl.

5 R<sup>6</sup> is an optionally substituted alkyl (including lower alkyl), cyano (CN), CH<sub>3</sub>, OCH<sub>3</sub>, OCH<sub>2</sub>CH<sub>3</sub>, hydroxy methyl (CH<sub>2</sub>OH), fluoromethyl (CH<sub>2</sub>F), azido (N<sub>3</sub>), CHCN, CH<sub>2</sub>N<sub>3</sub>, CH<sub>2</sub>NH<sub>2</sub>, CH<sub>2</sub>NHCH<sub>3</sub>, CH<sub>2</sub>N(CH<sub>3</sub>)<sub>2</sub>, alkyne (optionally substituted), or fluoro;

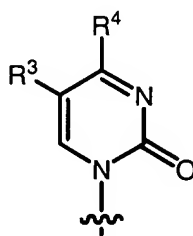
or its pharmaceutically acceptable salt or prodrug thereof, optionally in a pharmaceutically acceptable carrier.

10 100. A method for the treatment or prophylaxis of a Dengue virus infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside (β-D or β-L) of the formula:



15 wherein

Base is



20 R<sup>1</sup> and R<sup>7</sup> are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including

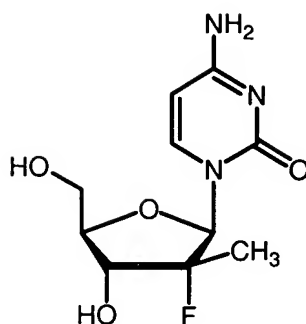
optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid, a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein R<sup>1</sup> or R<sup>7</sup> is independently H or phosphate; R<sup>1</sup> and R<sup>7</sup> can also be linked with cyclic phosphate group;

R<sup>3</sup> and R<sup>4</sup> are independently H, halogen including F, Cl, Br, I, OH, OR', SH, SR', NH<sub>2</sub>, NHR', NR'<sub>2</sub>, lower alkyl of C<sub>1</sub>-C<sub>6</sub>, halogenated (F, Cl, Br, I) lower alkyl of C<sub>1</sub>-C<sub>6</sub> such as CF<sub>3</sub> and CH<sub>2</sub>CH<sub>2</sub>F, lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CH<sub>2</sub>, halogenated (F, Cl, Br, I) lower alkenyl of C<sub>2</sub>-C<sub>6</sub> such as CH=CHCl, CH=CHBr and CH=CHI, lower alkynyl of C<sub>2</sub>-C<sub>6</sub> such as C≡CH, halogenated (F, Cl, Br, I) lower alkynyl of C<sub>2</sub>-C<sub>6</sub>, lower alkoxy of C<sub>1</sub>-C<sub>6</sub> such as CH<sub>2</sub>OH and CH<sub>2</sub>CH<sub>2</sub>OH, halogenated (F, Cl, Br, I) lower alkoxy of C<sub>1</sub>-C<sub>6</sub>, CO<sub>2</sub>H, CO<sub>2</sub>R', CONH<sub>2</sub>, CONHR', CONR'<sub>2</sub>, CH=CHCO<sub>2</sub>H, CH=CHCO<sub>2</sub>R';

R' is an optionally substituted alkyl of C<sub>1</sub>-C<sub>12</sub> (particularly when the alkyl is an amino acid residue), cycloalkyl, optionally substituted alkynyl of C<sub>2</sub>-C<sub>6</sub>, optionally substituted lower alkenyl of C<sub>2</sub>-C<sub>6</sub>, or optionally substituted acyl.

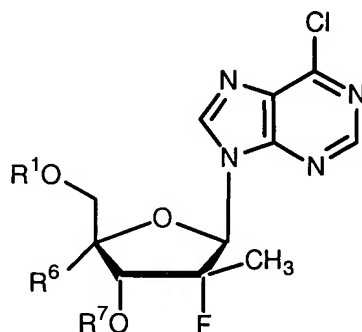
or its pharmaceutically acceptable salt or prodrug thereof, optionally in a pharmaceutically acceptable carrier.

101. A method for the treatment or prophylaxis of a Dengue virus infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside (β-D) or its pharmaceutically acceptable salt or prodrug thereof of the formula:



optionally in a pharmaceutically acceptable carrier.

102. A method for the treatment or prophylaxis of a Dengue virus infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside ( $\beta$ -D) or its pharmaceutically acceptable salt or prodrug thereof of the formula:



wherein

$R^1$  and  $R^7$  are independently H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid, a carbohydrate, a peptide, a cholesterol, or other pharmaceutically

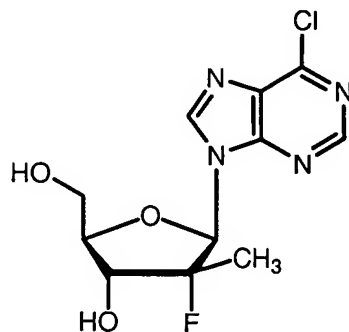
acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein  $R^1$  or  $R^7$  is independently H or phosphate;  $R^1$  and  $R^7$  can also be linked with cyclic phosphate group; and,

5  $R^6$  is an optionally substituted alkyl (including lower alkyl), cyano (CN),  $CH_3$ ,  $OCH_3$ ,  $OCH_2CH_3$ , hydroxy methyl ( $CH_2OH$ ), fluoromethyl ( $CH_2F$ ), azido ( $N_3$ ),  $CHCN$ ,  $CH_2N_3$ ,  $CH_2NH_2$ ,  $CH_2NHCH_3$ ,  $CH_2N(CH_3)_2$ , alkyne (optionally substituted), or fluoro and,

optionally in a pharmaceutically acceptable carrier.

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103. A method for the treatment or prophylaxis of a Dengue virus infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside ( $\beta$ -D) or its pharmaceutically acceptable salt or prodrug thereof of the formula:

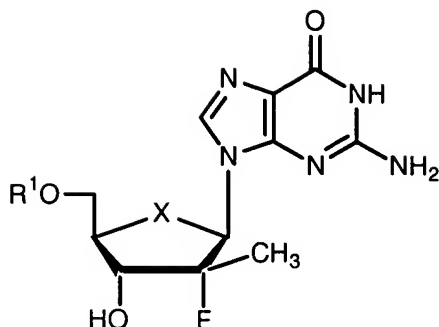


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and optionally a pharmaceutically acceptable carrier.

104. A method for the treatment or prophylaxis of a Dengue virus infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside ( $\beta$ -D) or its pharmaceutically acceptable salt or prodrug thereof of the formula:

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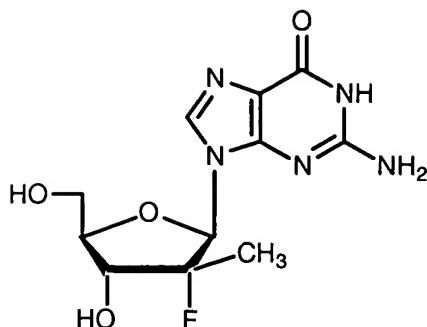


wherein

X is O, S, CH<sub>2</sub>, Se, NH, N-alkyl, CHW (*R*, *S*, or racemic), C(W)<sub>2</sub>, wherein W is F, Cl, Br, or I; and

5           R<sup>1</sup> is H, phosphate, including monophosphate, diphosphate, triphosphate, or a stabilized phosphate prodrug, H-phosphonate, including stabilized H-phosphonates, acyl, including optionally substituted phenyl and lower acyl, alkyl, including lower alkyl, O-substituted carboxyalkylamino or its peptide derivatives, sulfonate ester, including alkyl or arylalkyl  
10           sulfonyl, including methanesulfonyl and benzyl, wherein the phenyl group is optionally substituted, a lipid, including a phospholipid, an L or D-amino acid, a carbohydrate, a peptide, a cholesterol, or other pharmaceutically acceptable leaving group which when administered *in vivo* is capable of providing a compound wherein R<sup>1</sup> is H or phosphate;  
15           optionally in a pharmaceutically acceptable carrier.

105. A method for the treatment or prophylaxis of a Dengue virus infection comprising administering to a host an antivirally effective amount of a (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside (β-*D*) or its pharmaceutically acceptable salt or prodrug thereof  
20           of the formula:



optionally in a pharmaceutically acceptable carrier.

106. The method of 31, wherein the antivirally effective amount of (2'*R*)-2'-deoxy-  
 2'-fluoro-2'-*C*-methyl nucleoside is administered in combination or alternation with at least  
 one treatment selected from the group consisting of: interferon, including interferon alpha 2a,  
 interferon alpha 2b, a pegylated interferon, interferon beta, interferon gamma, interferon tau  
 and interferon omega; an interleukin, including interleukin 10 and interleukin 12; ribavirin;  
 interferon alpha or pegylated interferon alpha in combination with ribavirin or levovirin;  
 levovirin; a protease inhibitor including an NS3 inhibitor, a NS3-4A inhibitor; a helicase  
 inhibitor; a polymerase inhibitor including HCV RNA polymerase and NS5B polymerase  
 inhibitor; gliotoxin; an IRES inhibitor; and antisense oligonucleotide; a thiazolidine  
 derivative; a benzanilide, a ribozyme; another nucleoside, nucleoside prodrug or nucleoside  
 derivative; a 1-amino-alkylcyclohexane; an antioxidant including vitamin E; squalene;  
 amantadine; a bile acid; N-(phosphonoacetyl)-L-aspartic acid; a benzenedicarboxamide;  
 polyadenylic acid; a benzimidazoles; thymosin; a beta tubulin inhibitor; a prophylactic  
 vaccine; an immune modulator, an IMPDH inhibitor; silybin-phosphatidylcholine  
 phytosome; and mycophenolate.

107. The method of 41, wherein the antivirally effective amount of (2'*R*)-2'-deoxy-  
 2'-fluoro-2'-*C*-methyl nucleoside is administered in combination or alternation with at least  
 one treatment selected from the group consisting of: interferon, including interferon alpha 2a,  
 interferon alpha 2b, a pegylated interferon, interferon beta, interferon gamma, interferon tau  
 and interferon omega; an interleukin, including interleukin 10 and interleukin 12; ribavirin;

interferon alpha or pegylated interferon alpha in combination with ribavirin or levovirin; levovirin; a protease inhibitor including an NS3 inhibitor, a NS3-4A inhibitor; a helicase inhibitor; a polymerase inhibitor including HCV RNA polymerase and NS5B polymerase inhibitor; gliotoxin; an IRES inhibitor; and antisense oligonucleotide; a thiazolidine derivative; a benzanilide, a ribozyme; another nucleoside, nucleoside prodrug or nucleoside derivative; a 1-amino-alkylcyclohexane; an antioxidant including vitamin E; squalene; amantadine; a bile acid; N-(phosphonoacetyl)-L-aspartic acid; a benzenedicarboxamide; polyadenylic acid; a benzimidazoles; thymosin; a beta tubulin inhibitor; a prophylactic vaccine; an immune modulator, an IMPDH inhibitor; silybin-phosphatidylcholine phytosome; and mycophenolate.

108. The method of 43, wherein the antivirally effective amount of (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside is administered in combination or alternation with at least one treatment selected from the group consisting of: interferon, including interferon alpha 2a, interferon alpha 2b, a pegylated interferon, interferon beta, interferon gamma, interferon tau and interferon omega; an interleukin, including interleukin 10 and interleukin 12; ribavirin; interferon alpha or pegylated interferon alpha in combination with ribavirin or levovirin; levovirin; a protease inhibitor including an NS3 inhibitor, a NS3-4A inhibitor; a helicase inhibitor; a polymerase inhibitor including HCV RNA polymerase and NS5B polymerase inhibitor; gliotoxin; an IRES inhibitor; and antisense oligonucleotide; a thiazolidine derivative; a benzanilide, a ribozyme; another nucleoside, nucleoside prodrug or nucleoside derivative; a 1-amino-alkylcyclohexane; an antioxidant including vitamin E; squalene; amantadine; a bile acid; N-(phosphonoacetyl)-L-aspartic acid; a benzenedicarboxamide; polyadenylic acid; a benzimidazoles; thymosin; a beta tubulin inhibitor; a prophylactic vaccine; an immune modulator, an IMPDH inhibitor; silybin-phosphatidylcholine phytosome; and mycophenolate.

109. The method of 45, wherein the antivirally effective amount of (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside is administered in combination or alternation with at least one treatment selected from the group consisting of: interferon, including interferon alpha 2a,



interferon alpha 2b, a pegylated interferon, interferon beta, interferon gamma, interferon tau and interferon omega; an interleukin, including interleukin 10 and interleukin 12; ribavirin; interferon alpha or pegylated interferon alpha in combination with ribavirin or levovirin; levovirin; a protease inhibitor including an NS3 inhibitor, a NS3-4A inhibitor; a helicase inhibitor; a polymerase inhibitor including HCV RNA polymerase and NS5B polymerase inhibitor; gliotoxin; an IRES inhibitor; and antisense oligonucleotide; a thiazolidine derivative; a benzanilide, a ribozyme; another nucleoside, nucleoside prodrug or nucleoside derivative; a 1-amino-alkylcyclohexane; an antioxidant including vitamin E; squalene; amantadine; a bile acid; N-(phosphonoacetyl)-L-aspartic acid; a benzenedicarboxamide; polyadenylic acid; a benzimidazoles; thymosin; a beta tubulin inhibitor; a prophylactic vaccine; an immune modulator, an IMPDH inhibitor; silybin-phosphatidylcholine phytosome; and mycophenolate.

110. The method of 46, wherein the antivirally effective amount of (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside is administered in combination or alternation with at least one treatment selected from the group consisting of: interferon, including interferon alpha 2a, interferon alpha 2b, a pegylated interferon, interferon beta, interferon gamma, interferon tau and interferon omega; an interleukin, including interleukin 10 and interleukin 12; ribavirin; interferon alpha or pegylated interferon alpha in combination with ribavirin or levovirin; levovirin; a protease inhibitor including an NS3 inhibitor, a NS3-4A inhibitor; a helicase inhibitor; a polymerase inhibitor including HCV RNA polymerase and NS5B polymerase inhibitor; gliotoxin; an IRES inhibitor; and antisense oligonucleotide; a thiazolidine derivative; a benzanilide, a ribozyme; another nucleoside, nucleoside prodrug or nucleoside derivative; a 1-amino-alkylcyclohexane; an antioxidant including vitamin E; squalene; amantadine; a bile acid; N-(phosphonoacetyl)-L-aspartic acid; a benzenedicarboxamide; polyadenylic acid; a benzimidazoles; thymosin; a beta tubulin inhibitor; a prophylactic vaccine; an immune modulator, an IMPDH inhibitor; silybin-phosphatidylcholine phytosome; and mycophenolate.

111. The method of 56, wherein the antivirally effective amount of (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside is administered in combination or alternation with at least one treatment selected from the group consisting of: interferon, including interferon alpha 2a, interferon alpha 2b, a pegylated interferon, interferon beta, interferon gamma, interferon tau and interferon omega; an interleukin, including interleukin 10 and interleukin 12; ribavirin; interferon alpha or pegylated interferon alpha in combination with ribavirin or levovirin; levovirin; a protease inhibitor including an NS3 inhibitor, a NS3-4A inhibitor; a helicase inhibitor; a polymerase inhibitor including HCV RNA polymerase and NS5B polymerase inhibitor; gliotoxin; an IRES inhibitor; and antisense oligonucleotide; a thiazolidine derivative; a benzanilide, a ribozyme; another nucleoside, nucleoside prodrug or nucleoside derivative; a 1-amino-alkylcyclohexane; an antioxidant including vitamin E; squalene; amantadine; a bile acid; N-(phosphonoacetyl)-L-aspartic acid; a benzenedicarboxamide; polyadenylic acid; a benzimidazoles; thymosin; a beta tubulin inhibitor; a prophylactic vaccine; an immune modulator, an IMPDH inhibitor; silybin-phosphatidylcholine phytosome; and mycophenolate.

112. The method of 58, wherein the antivirally effective amount of (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside is administered in combination or alternation with at least one treatment selected from the group consisting of: interferon, including interferon alpha 2a, interferon alpha 2b, a pegylated interferon, interferon beta, interferon gamma, interferon tau and interferon omega; an interleukin, including interleukin 10 and interleukin 12; ribavirin; interferon alpha or pegylated interferon alpha in combination with ribavirin or levovirin; levovirin; a protease inhibitor including an NS3 inhibitor, a NS3-4A inhibitor; a helicase inhibitor; a polymerase inhibitor including HCV RNA polymerase and NS5B polymerase inhibitor; gliotoxin; an IRES inhibitor; and antisense oligonucleotide; a thiazolidine derivative; a benzanilide, a ribozyme; another nucleoside, nucleoside prodrug or nucleoside derivative; a 1-amino-alkylcyclohexane; an antioxidant including vitamin E; squalene; amantadine; a bile acid; N-(phosphonoacetyl)-L-aspartic acid; a benzenedicarboxamide; polyadenylic acid; a benzimidazoles; thymosin; a beta tubulin inhibitor; a prophylactic

vaccine; an immune modulator, an IMPDH inhibitor; silybin-phosphatidylcholine phytosome; and mycophenolate.

113. The method of 60, wherein the antivirally effective amount of (2'R)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside is administered in combination or alternation with at least one treatment selected from the group consisting of: interferon, including interferon alpha 2a, interferon alpha 2b, a pegylated interferon, interferon beta, interferon gamma, interferon tau and interferon omega; an interleukin, including interleukin 10 and interleukin 12; ribavirin; interferon alpha or pegylated interferon alpha in combination with ribavirin or levovirin; levovirin; a protease inhibitor including an NS3 inhibitor, a NS3-4A inhibitor; a helicase inhibitor; a polymerase inhibitor including HCV RNA polymerase and NS5B polymerase inhibitor; gliotoxin; an IRES inhibitor; and antisense oligonucleotide; a thiazolidine derivative; a benzanilide, a ribozyme; another nucleoside, nucleoside prodrug or nucleoside derivative; a 1-amino-alkylcyclohexane; an antioxidant including vitamin E; squalene; amantadine; a bile acid; N-(phosphonoacetyl)-L-aspartic acid; a benzenedicarboxamide; polyadenylic acid; a benzimidazoles; thymosin; a beta tubulin inhibitor; a prophylactic vaccine; an immune modulator, an IMPDH inhibitor; silybin-phosphatidylcholine phytosome; and mycophenolate.

114. The method of 61, wherein the antivirally effective amount of (2'R)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside is administered in combination or alternation with at least one treatment selected from the group consisting of: interferon, including interferon alpha 2a, interferon alpha 2b, a pegylated interferon, interferon beta, interferon gamma, interferon tau and interferon omega; an interleukin, including interleukin 10 and interleukin 12; ribavirin; interferon alpha or pegylated interferon alpha in combination with ribavirin or levovirin; levovirin; a protease inhibitor including an NS3 inhibitor, a NS3-4A inhibitor; a helicase inhibitor; a polymerase inhibitor including HCV RNA polymerase and NS5B polymerase inhibitor; gliotoxin; an IRES inhibitor; and antisense oligonucleotide; a thiazolidine derivative; a benzanilide, a ribozyme; another nucleoside, nucleoside prodrug or nucleoside derivative; a 1-amino-alkylcyclohexane; an antioxidant including vitamin E; squalene;

amantadine; a bile acid; N-(phosphonoacetyl)-L-aspartic acid; a benzenedicarboxamide; polyadenylic acid; a benzimidazoles; thymosin; a beta tubulin inhibitor; a prophylactic vaccine; an immune modulator, an IMPDH inhibitor; silybin-phosphatidylcholine phytosome; and mycophenolate.

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115. The method of 71, wherein the antivirally effective amount of (2'R)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside is administered in combination or alternation with at least one treatment selected from the group consisting of: interferon, including interferon alpha 2a, interferon alpha 2b, a pegylated interferon, interferon beta, interferon gamma, interferon tau and interferon omega; an interleukin, including interleukin 10 and interleukin 12; ribavirin; interferon alpha or pegylated interferon alpha in combination with ribavirin or levovirin; levovirin; a protease inhibitor including an NS3 inhibitor, a NS3-4A inhibitor; a helicase inhibitor; a polymerase inhibitor including HCV RNA polymerase and NS5B polymerase inhibitor; gliotoxin; an IRES inhibitor; and antisense oligonucleotide; a thiazolidine derivative; a benzanilide, a ribozyme; another nucleoside, nucleoside prodrug or nucleoside derivative; a 1-amino-alkylcyclohexane; an antioxidant including vitamin E; squalene; amantadine; a bile acid; N-(phosphonoacetyl)-L-aspartic acid; a benzenedicarboxamide; polyadenylic acid; a benzimidazoles; thymosin; a beta tubulin inhibitor; a prophylactic vaccine; an immune modulator, an IMPDH inhibitor; silybin-phosphatidylcholine phytosome; and mycophenolate.

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116. The method of 73, wherein the antivirally effective amount of (2'R)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside is administered in combination or alternation with at least one treatment selected from the group consisting of: interferon, including interferon alpha 2a, interferon alpha 2b, a pegylated interferon, interferon beta, interferon gamma, interferon tau and interferon omega; an interleukin, including interleukin 10 and interleukin 12; ribavirin; interferon alpha or pegylated interferon alpha in combination with ribavirin or levovirin; levovirin; a protease inhibitor including an NS3 inhibitor, a NS3-4A inhibitor; a helicase inhibitor; a polymerase inhibitor including HCV RNA polymerase and NS5B polymerase inhibitor; gliotoxin; an IRES inhibitor; and antisense oligonucleotide; a thiazolidine

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derivative; a benzanilide, a ribozyme; another nucleoside, nucleoside prodrug or nucleoside derivative; a 1-amino-alkylcyclohexane; an antioxidant including vitamin E; squalene; amantadine; a bile acid; N-(phosphonoacetyl)-L-aspartic acid; a benzenedicarboxamide; polyadenylic acid; a benzimidazoles; thymosin; a beta tubulin inhibitor; a prophylactic vaccine; an immune modulator, an IMPDH inhibitor; silybin-phosphatidylcholine phytosome; and mycophenolate.

117. The method of 75, wherein the antivirally effective amount of (2'R)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside is administered in combination or alternation with at least one treatment selected from the group consisting of: interferon, including interferon alpha 2a, interferon alpha 2b, a pegylated interferon, interferon beta, interferon gamma, interferon tau and interferon omega; an interleukin, including interleukin 10 and interleukin 12; ribavirin; interferon alpha or pegylated interferon alpha in combination with ribavirin or levovirin; levovirin; a protease inhibitor including an NS3 inhibitor, a NS3-4A inhibitor; a helicase inhibitor; a polymerase inhibitor including HCV RNA polymerase and NS5B polymerase inhibitor; gliotoxin; an IRES inhibitor; and antisense oligonucleotide; a thiazolidine derivative; a benzanilide, a ribozyme; another nucleoside, nucleoside prodrug or nucleoside derivative; a 1-amino-alkylcyclohexane; an antioxidant including vitamin E; squalene; amantadine; a bile acid; N-(phosphonoacetyl)-L-aspartic acid; a benzenedicarboxamide; polyadenylic acid; a benzimidazoles; thymosin; a beta tubulin inhibitor; a prophylactic vaccine; an immune modulator, an IMPDH inhibitor; silybin-phosphatidylcholine phytosome; and mycophenolate.

118. The method of 76, wherein the antivirally effective amount of (2'R)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside is administered in combination or alternation with at least one treatment selected from the group consisting of: interferon, including interferon alpha 2a, interferon alpha 2b, a pegylated interferon, interferon beta, interferon gamma, interferon tau and interferon omega; an interleukin, including interleukin 10 and interleukin 12; ribavirin; interferon alpha or pegylated interferon alpha in combination with ribavirin or levovirin; levovirin; a protease inhibitor including an NS3 inhibitor, a NS3-4A inhibitor; a helicase

inhibitor; a polymerase inhibitor including HCV RNA polymerase and NS5B polymerase inhibitor; gliotoxin; an IRES inhibitor; and antisense oligonucleotide; a thiazolidine derivative; a benzanilide, a ribozyme; another nucleoside, nucleoside prodrug or nucleoside derivative; a 1-amino-alkylcyclohexane; an antioxidant including vitamin E; squalene; amantadine; a bile acid; N-(phosphonoacetyl)-L-aspartic acid; a benzenedicarboxamide; polyadenylic acid; a benzimidazoles; thymosin; a beta tubulin inhibitor; a prophylactic vaccine; an immune modulator, an IMPDH inhibitor; silybin-phosphatidylcholine phytosome; and mycophenolate.

119. The method of 86, wherein the antivirally effective amount of (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside is administered in combination or alternation with at least one treatment selected from the group consisting of: interferon, including interferon alpha 2a, interferon alpha 2b, a pegylated interferon, interferon beta, interferon gamma, interferon tau and interferon omega; an interleukin, including interleukin 10 and interleukin 12; ribavirin; interferon alpha or pegylated interferon alpha in combination with ribavirin or levovirin; levovirin; a protease inhibitor including an NS3 inhibitor, a NS3-4A inhibitor; a helicase inhibitor; a polymerase inhibitor including HCV RNA polymerase and NS5B polymerase inhibitor; gliotoxin; an IRES inhibitor; and antisense oligonucleotide; a thiazolidine derivative; a benzanilide, a ribozyme; another nucleoside, nucleoside prodrug or nucleoside derivative; a 1-amino-alkylcyclohexane; an antioxidant including vitamin E; squalene; amantadine; a bile acid; N-(phosphonoacetyl)-L-aspartic acid; a benzenedicarboxamide; polyadenylic acid; a benzimidazoles; thymosin; a beta tubulin inhibitor; a prophylactic vaccine; an immune modulator, an IMPDH inhibitor; silybin-phosphatidylcholine phytosome; and mycophenolate.

120. The method of 88, wherein the antivirally effective amount of (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside is administered in combination or alternation with at least one treatment selected from the group consisting of: interferon, including interferon alpha 2a, interferon alpha 2b, a pegylated interferon, interferon beta, interferon gamma, interferon tau and interferon omega; an interleukin, including interleukin 10 and interleukin 12; ribavirin;

interferon alpha or pegylated interferon alpha in combination with ribavirin or levovirin; levovirin; a protease inhibitor including an NS3 inhibitor, a NS3-4A inhibitor; a helicase inhibitor; a polymerase inhibitor including HCV RNA polymerase and NS5B polymerase inhibitor; gliotoxin; an IRES inhibitor; and antisense oligonucleotide; a thiazolidine derivative; a benzanilide, a ribozyme; another nucleoside, nucleoside prodrug or nucleoside derivative; a 1-amino-alkylcyclohexane; an antioxidant including vitamin E; squalene; amantadine; a bile acid; N-(phosphonoacetyl)-L-aspartic acid; a benzenedicarboxamide; polyadenylic acid; a benzimidazoles; thymosin; a beta tubulin inhibitor; a prophylactic vaccine; an immune modulator, an IMPDH inhibitor; silybin-phosphatidylcholine phytosome; and mycophenolate.

121. The method of 90, wherein the antivirally effective amount of (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside is administered in combination or alternation with at least one treatment selected from the group consisting of: interferon, including interferon alpha 2a, interferon alpha 2b, a pegylated interferon, interferon beta, interferon gamma, interferon tau and interferon omega; an interleukin, including interleukin 10 and interleukin 12; ribavirin; interferon alpha or pegylated interferon alpha in combination with ribavirin or levovirin; levovirin; a protease inhibitor including an NS3 inhibitor, a NS3-4A inhibitor; a helicase inhibitor; a polymerase inhibitor including HCV RNA polymerase and NS5B polymerase inhibitor; gliotoxin; an IRES inhibitor; and antisense oligonucleotide; a thiazolidine derivative; a benzanilide, a ribozyme; another nucleoside, nucleoside prodrug or nucleoside derivative; a 1-amino-alkylcyclohexane; an antioxidant including vitamin E; squalene; amantadine; a bile acid; N-(phosphonoacetyl)-L-aspartic acid; a benzenedicarboxamide; polyadenylic acid; a benzimidazoles; thymosin; a beta tubulin inhibitor; a prophylactic vaccine; an immune modulator, an IMPDH inhibitor; silybin-phosphatidylcholine phytosome; and mycophenolate.

122. The method of 91, wherein the antivirally effective amount of (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside is administered in combination or alternation with at least one treatment selected from the group consisting of: interferon, including interferon alpha 2a,

interferon alpha 2b, a pegylated interferon, interferon beta, interferon gamma, interferon tau and interferon omega; an interleukin, including interleukin 10 and interleukin 12; ribavirin; interferon alpha or pegylated interferon alpha in combination with ribavirin or levovirin; levovirin; a protease inhibitor including an NS3 inhibitor, a NS3-4A inhibitor; a helicase inhibitor; a polymerase inhibitor including HCV RNA polymerase and NS5B polymerase inhibitor; gliotoxin; an IRES inhibitor; and antisense oligonucleotide; a thiazolidine derivative; a benzanilide, a ribozyme; another nucleoside, nucleoside prodrug or nucleoside derivative; a 1-amino-alkylcyclohexane; an antioxidant including vitamin E; squalene; amantadine; a bile acid; N-(phosphonoacetyl)-L-aspartic acid; a benzenedicarboxamide; polyadenylic acid; a benzimidazoles; thymosin; a beta tubulin inhibitor; a prophylactic vaccine; an immune modulator, an IMPDH inhibitor; silybin-phosphatidylcholine phytosome; and mycophenolate.

123. The method of 101, wherein the antivirally effective amount of (2'R)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside is administered in combination or alternation with at least one treatment selected from the group consisting of: interferon, including interferon alpha 2a, interferon alpha 2b, a pegylated interferon, interferon beta, interferon gamma, interferon tau and interferon omega; an interleukin, including interleukin 10 and interleukin 12; ribavirin; interferon alpha or pegylated interferon alpha in combination with ribavirin or levovirin; levovirin; a protease inhibitor including an NS3 inhibitor, a NS3-4A inhibitor; a helicase inhibitor; a polymerase inhibitor including HCV RNA polymerase and NS5B polymerase inhibitor; gliotoxin; an IRES inhibitor; and antisense oligonucleotide; a thiazolidine derivative; a benzanilide, a ribozyme; another nucleoside, nucleoside prodrug or nucleoside derivative; a 1-amino-alkylcyclohexane; an antioxidant including vitamin E; squalene; amantadine; a bile acid; N-(phosphonoacetyl)-L-aspartic acid; a benzenedicarboxamide; polyadenylic acid; a benzimidazoles; thymosin; a beta tubulin inhibitor; a prophylactic vaccine; an immune modulator, an IMPDH inhibitor; silybin-phosphatidylcholine phytosome; and mycophenolate.

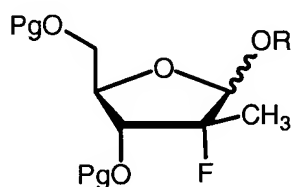


124. The method of 103, wherein the antivirally effective amount of (2'R)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside is administered in combination or alternation with at least one treatment selected from the group consisting of: interferon, including interferon alpha 2a, interferon alpha 2b, a pegylated interferon, interferon beta, interferon gamma, interferon tau and interferon omega; an interleukin, including interleukin 10 and interleukin 12; ribavirin; interferon alpha or pegylated interferon alpha in combination with ribavirin or levovirin; levovirin; a protease inhibitor including an NS3 inhibitor, a NS3-4A inhibitor; a helicase inhibitor; a polymerase inhibitor including HCV RNA polymerase and NS5B polymerase inhibitor; gliotoxin; an IRES inhibitor; and antisense oligonucleotide; a thiazolidine derivative; a benzanilide, a ribozyme; another nucleoside, nucleoside prodrug or nucleoside derivative; a 1-amino-alkylcyclohexane; an antioxidant including vitamin E; squalene; amantadine; a bile acid; N-(phosphonoacetyl)-L-aspartic acid; a benzenedicarboxamide; polyadenylic acid; a benzimidazoles; thymosin; a beta tubulin inhibitor; a prophylactic vaccine; an immune modulator, an IMPDH inhibitor; silybin-phosphatidylcholine phytosome; and mycophenolate.

125. The method of 105, wherein the antivirally effective amount of (2'R)-2'-deoxy-2'-fluoro-2'-C-methyl nucleoside is administered in combination or alternation with at least one treatment selected from the group consisting of: interferon, including interferon alpha 2a, interferon alpha 2b, a pegylated interferon, interferon beta, interferon gamma, interferon tau and interferon omega; an interleukin, including interleukin 10 and interleukin 12; ribavirin; interferon alpha or pegylated interferon alpha in combination with ribavirin or levovirin; levovirin; a protease inhibitor including an NS3 inhibitor, a NS3-4A inhibitor; a helicase inhibitor; a polymerase inhibitor including HCV RNA polymerase and NS5B polymerase inhibitor; gliotoxin; an IRES inhibitor; and antisense oligonucleotide; a thiazolidine derivative; a benzanilide, a ribozyme; another nucleoside, nucleoside prodrug or nucleoside derivative; a 1-amino-alkylcyclohexane; an antioxidant including vitamin E; squalene; amantadine; a bile acid; N-(phosphonoacetyl)-L-aspartic acid; a benzenedicarboxamide; polyadenylic acid; a benzimidazoles; thymosin; a beta tubulin

inhibitor; a prophylactic vaccine; an immune modulator, an IMPDH inhibitor; silybin-phosphatidylcholine phytosome; and mycophenolate.

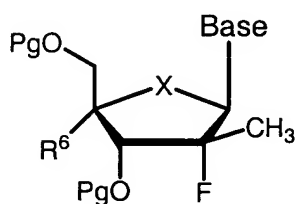
126. A method of synthesizing a (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside (β-D or β-L) comprising glycosylation of a nucleobase with an intermediate structure:



1-4

wherein R is lower alkyl, acyl, benzoyl, or mesyl; and Pg is any acceptable protecting group consisting of but not limited to C(O)-alkyl, C(O)Ph, C(O)aryl, CH<sub>3</sub>, CH<sub>2</sub>-alkyl, CH<sub>2</sub>-alkenyl, CH<sub>2</sub>Ph, CH<sub>2</sub>-aryl, CH<sub>2</sub>O-alkyl, CH<sub>2</sub>O-aryl, SO<sub>2</sub>-alkyl, SO<sub>2</sub>-aryl, *tert*-butyldimethylsilyl, *tert*-butyldiphenylsilyl, or both Pg's may come together to form a 1,3-(1,1,3,3-tetraisopropylidisiloxanylidene).

127. A method of synthesizing a (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside (β-D or β-L) comprising selective deprotection of either Pg in an intermediate of the structure:



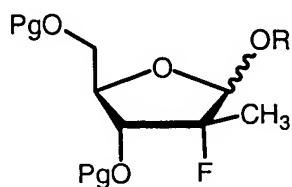
2-5

wherein, X is O, S, CH<sub>2</sub>, Se, NH, N-alkyl, CHW (*R*, *S*, or racemic), C(W)<sub>2</sub>, wherein W is F, Cl, Br, or I; and Pg is independently any pharmaceutically acceptable protecting group

selected from the group consisting of C(O)-alkyl, C(O)Ph, C(O)aryl, CH<sub>3</sub>, CH<sub>2</sub>-alkyl, CH<sub>2</sub>-alkenyl, CH<sub>2</sub>Ph, CH<sub>2</sub>-aryl, CH<sub>2</sub>O-alkyl, CH<sub>2</sub>O-aryl, SO<sub>2</sub>-alkyl, SO<sub>2</sub>-aryl, *tert*-butyldimethylsilyl, *tert*-butyldiphenylsilyl, or both Pg's may come together to form a 1,3-(1,1,3,3-tetraisopropylidisiloxanylidene).

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128. An intermediate in the synthesis of a (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside ( $\beta$ -D or  $\beta$ -L), wherein the intermediate is of the structure:

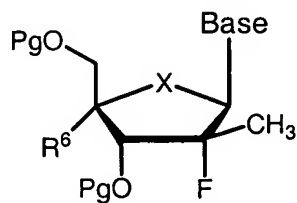


1-4

10 wherein R is lower alkyl, acyl, benzoyl, or mesyl; and Pg is any acceptable protecting group consisting of but not limited to C(O)-alkyl, C(O)Ph, C(O)aryl, CH<sub>3</sub>, CH<sub>2</sub>-alkyl, CH<sub>2</sub>-alkenyl, CH<sub>2</sub>Ph, CH<sub>2</sub>-aryl, CH<sub>2</sub>O-alkyl, CH<sub>2</sub>O-aryl, SO<sub>2</sub>-alkyl, SO<sub>2</sub>-aryl, *tert*-butyldimethylsilyl, *tert*-butyldiphenylsilyl, or both Pg's may come together to form a 1,3-(1,1,3,3-tetraisopropylidisiloxanylidene).

15

129. An intermediate in the synthesis of a (2'*R*)-2'-deoxy-2'-fluoro-2'-*C*-methyl nucleoside ( $\beta$ -D or  $\beta$ -L), wherein the intermediate is of the structure:



2-5

wherein, X is O, S, CH<sub>2</sub>, Se, NH, N-alkyl, CHW (*R*, *S*, or racemic), C(W)<sub>2</sub>, wherein W is F, Cl, Br, or I; and Pg is independently any pharmaceutically acceptable protecting group selected from the group consisting of C(O)-alkyl, C(O)Ph, C(O)aryl, CH<sub>3</sub>, CH<sub>2</sub>-alkyl, CH<sub>2</sub>-alkenyl, CH<sub>2</sub>Ph, CH<sub>2</sub>-aryl, CH<sub>2</sub>O-alkyl, CH<sub>2</sub>O-aryl, SO<sub>2</sub>-alkyl, SO<sub>2</sub>-aryl, *tert*-butyldimethylsilyl, *tert*-butyldiphenylsilyl, or both Pg's may come together to form a 1,3-(1,1,3,3-tetraisopropylidisiloxanylidene).